

chain nodes:

7 8 9 10 11 13 15 17 18

ring nodes:

1 2 3 4 5 6

chain bonds:

2-7 3-17 5-8 7-15 8-9 9-10 10-11 10-13 17-18

ring bonds:

1-2 1-6 2-3 3-4 4-5 5-6

exact/norm bonds:

2-7 3-17 7-15 10-11 10-13 17-18

exact bonds:

5-8 8-9 9-10

normalized bonds:

1-2 1-6 2-3 3-4 4-5 5-6

G1:OH,N

G2:H,Cb,Cy,Hy,Ak

Match level:

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS8:CLASS9:CLASS10:CLASS11:CLASS13:CLASS 15:CLASS17:CLASS18:CLASS

(FILE 'HOME' ENTERED AT 10:56:25 ON 08 FEB 2007)

```
FILE 'REGISTRY' ENTERED AT 11:04:58 ON 08 FEB 2007
                STRUCTURE UPLOADED
L1
           7462 S L1 SSS FULL
L2
     FILE 'CAPLUS, USPATFULL' ENTERED AT 11:05:33 ON 08 FEB 2007
            853 S L2 AND (HYPERTENSION OR HYPERTENSIVE OR HIGH BLOOD PRESSURE
L3
            510 S L3 AND (HYPERTENSION OR HIGH BLOOD PRESSURE)
L4
            611 S L3 AND (HYPERTENSION OR HIGH BLOOD PRESSURE OR ANTIHYPERTENSI
L5
            586 DUP REM L5 (25 DUPLICATES REMOVED)
L6
            571 S L6 NOT FERULIC ACID
L7
            571 FOCUS L7 1-
L8
L9
            382 S L8 AND PD <= 2001
            382 FOCUS L9 1-
L10
=> s 110 not caffeic acid
          378 L10 NOT CAFFEIC ACID
=> s l11 and (hypertension or high blood pressure)
           309 L11 AND (HYPERTENSION OR HIGH BLOOD PRESSURE)
L12
```

=> d ibib abs hitstr 51-100

L26 ANSWER 48 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1976:539727 CAPLUS

DOCUMENT NUMBER: 85:139727

TITLE: Isolation and synthesis of pinoresinol diglucoside, a

major antihypertensive principle of Tu-Chung (Eucommia

ulmoides, Oliver)

AUTHOR(S): Sih, Charles J.; Ravikumar, P. R.; Huang, Fu-Chih;

Buckner, Carl; Whitlock, Howard, Jr.

CORPORATE SOURCE: Sch. Pharm., Univ. Wisconsin, Madison, WI, USA

SOURCE: Journal of the American Chemical Society (1976

), 98(17), 5412-13

CODEN: JACSAT; ISSN: 0002-7863

DOCUMENT TYPE: Journal LANGUAGE: English

AB The major antihypertensive principle of Tu-Chung bark (E. ulmoides) was isolated and shown to be pinoresinol-di- β -D-glucoside. Exposure of

coniferyl alc. to the chloroperoxidase-containing microorganism, Caldariomyces

fumago, gave (±)-pinoresinol and (±)-cis-dehydrodiconiferyl alc.

The resulting (\pm) -pinoresinol was reacted with 2 moles of α -bromoacetoglucose to yield a product whose antihypertensive activity was indistinguishable from the natural product.

IT 458-35-5

RL: RCT (Reactant); RACT (Reactant or reagent)
 (dimerization of)

RN 458-35-5 CAPLUS

CN Phenol, 4-(3-hydroxy-1-propenyl)-2-methoxy- (9CI) (CA INDEX NAME)

$$CH = CH - CH_2 - OH$$

OMe

L26 ANSWER 46 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1978:87653 CAPLUS

DOCUMENT NUMBER: 88:87653

TITLE: Glycosides of 2,6-bis(hydroxyphenyl)-3,7-dioxabicyclo

[3.3.0] octane Sih, Charles John

PATENT ASSIGNEE(S): Wisconsin Alumni Research Foundation, USA

SOURCE: Ger. Offen., 25 pp.

CODEN: GWXXBX

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

INVENTOR(S):

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|------|----------|------------------|------------|
| | | | | |
| DE 2722911 | A1 | 19771201 | DE 1977-2722911 | 19770520 < |
| US 4103006 | Α | 19780725 | US 1976-688275 | 19760520 < |
| CA 1085826 | A1 | 19800916 | CA 1977-278165 | 19770511 < |
| GB 1575439 | Α | 19800924 | GB 1977-21229 | 19770519 < |
| FR 2351992 | A1 | 19771216 | FR 1977-15485 | 19770520 < |
| FR 2351992 | B1 | 19800411 | | |
| JP 53007698 | A | 19780124 | JP 1977-57799 | 19770520 < |
| PRIORITY APPLN. INFO.: | | | US 1976-688275 A | 19760520 |
| GI | | | | |

AB Compds. of structure I, where X1, X2, X3, and X4 are H, OH, Cl, NH2, lower alkyl, or lower alkoxy groups and R and R1 are mono- or disaccharides, are prepared by chemical synthesis, isolation from the tree Eucommia ulmoides, or by fermentation with Caldariomyces fumago. These compds. have antihypertensive properties. Thus, C. fumago was inoculated into flasks containing 500 mL medium composed of soybean meal 5, dextrose 20, NaCl 5, K2HPO4 5, and yeast extract 5 g/L. After 48 h at 25° with shaking, 2.5 g coniferyl alc. [458-35-5] dissolved in 4 mL DMF were added to each flask. After 16 addnl. h, the supernatant from 3 flasks were combined, acidified to pH 2.5, and extracted with EtOAc. The extract was evaporated and the oily residue

chromatographed on silica gel to yield 810 mg (\pm)-pinoresinol [4263-88-1] and 798 mg cis-dehydrodiconiferyl alc. [60536-58-5]. Pinoresinol diglucoside [63902-38-5] was prepared in a 34% yield by reaction of pinoresinol with Ag2O and aceto- α -bromoglucose [572-09-8].

IT 458-35-5

RL: BIOL (Biological study)

(pinoresinol manufacture from, with Caldariomyces fumago)

RN 458-35-5 CAPLUS

CN Phenol, 4-(3-hydroxy-1-propenyl)-2-methoxy- (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{CH-CH}_2\text{-OH} \\ \\ \text{OMe} \end{array}$$

L26 ANSWER 47 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1977:137539 CAPLUS

DOCUMENT NUMBER: 86:137539

Increased vesicular transfer of horseradish peroxidase TITLE:

across cerebral endothelium, evoked by acute

hypertension

Westergaard, E.; Van Deurs, B.; Broendsted, H. E. AUTHOR (S):

Anat. Dep. C, Univ. Copenhagen, Copenhagen, Den. CORPORATE SOURCE:

Acta Neuropathologica (1977), 37(2), 141-52 SOURCE:

CODEN: ANPTAL; ISSN: 0001-6322

DOCUMENT TYPE: Journal English LANGUAGE:

The permeability to i.v. injected horseradish peroxidase (HRP) was increased across the cerebral arterioles, capillaries and venules in acute induced hypertension. The tight junctions between endothelial cells were intact and prevented intercellular movement of peroxidase. Many HRP-labeled vesicles within the endothelial cells or connected with the luminal or abluminal surface, occurred in segments of the microvasculature. Otherwise the endothelium was unchanged. Thus, acute hypertension increases the vesicular transport of HRP across the

endothelium of cerebral arterioles, venules, and capillaries that normally

occurs to a small extent only after i.v. injection of the tracer.

IT 9003-99-0

RL: BIOL (Biological study)

(cerebral endothelium permeability to, in hypertension)

RN9003-99-0 CAPLUS

CN Peroxidase (9CI) (CA INDEX NAME) L26 ANSWER 34 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1995:514068 CAPLUS

DOCUMENT NUMBER: 122:281796

TITLE: Antihypertensive activity of phenolics from the flower

of Lonicera japonica

AUTHOR(S): Cheng, Juei-Tang; Lee, Yung-Yung; Hsu, Feng-Lin;

Chang, Wen; Niu, Chiang-Shan

CORPORATE SOURCE: College of Medicine, National Cheng Kung University,

Tainan, 70101, Taiwan

SOURCE: Chinese Pharmaceutical Journal (Taipei, Taiwan) (

1994), 46(6), 575-82

CODEN: CPHJEP; ISSN: 1016-1015

DOCUMENT TYPE: Journal LANGUAGE: English

The effects on blood pressure of phenolic compds. obtained from dried flower of Lonicera japonica Thunb. (Caprifoliaceae) were investigated in spontaneously hypertensive rats. Protocatechuic acid and Me caffeate were identified as the major active substances. Chlorogenic acid and five caffeoylquinic acids at higher doses possessed the delay hypotensive effect. Also, at the highest ED, all of these compds. except the

flavonoids rutin and luteolin produced hypotension in normotensive rats.

IT 327-97-9, Chlorogenic acid 2450-53-5 14534-61-3 29708-87-0, Methyl chlorogenate 141545-93-9

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)

(antihypertensive activity of phenolics from flower of Lonicera japonica)

RN 327-97-9 CAPLUS

CN Cyclohexanecarboxylic acid, 3-[[3-(3,4-dihydroxyphenyl)-1-oxo-2-propenyl]oxy]-1,4,5-trihydroxy-, (1S,3R,4R,5R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

RN 2450-53-5 CAPLUS

CN Cyclohexanecarboxylic acid, 3,5-bis[[3-(3,4-dihydroxyphenyl)-1-oxo-2-propenyl]oxy]-1,4-dihydroxy-, $(1\alpha,3R,4\alpha,5R)$ - (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

RN 14534-61-3 CAPLUS

CN Cyclohexanecarboxylic acid, 3,4-bis[[3-(3,4-dihydroxyphenyl)-1-oxo-2-propenyl]oxy]-1,5-dihydroxy-, (1S,3R,4R,5R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

RN 29708-87-0 CAPLUS

CN Cyclohexanecarboxylic acid, 3-[[3-(3,4-dihydroxyphenyl)-1-oxo-2-propenyl]oxy]-1,4,5-trihydroxy-, methyl ester, (1S,3R,4R,5R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

RN 141545-93-9 CAPLUS

CN Cyclohexanecarboxylic acid, 3,5-bis[[3-(3,4-dihydroxyphenyl)-1-oxo-2-

propenyl]oxy]-1,4-dihydroxy-, methyl ester, $(1\alpha,3R,4\alpha,5R)$ -(9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

L26 ANSWER 35 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1994:698944 CAPLUS

DOCUMENT NUMBER: 121:298944

TITLE: Immune complex glomerulonephritis is induced in rats

immunized with heterologous myeloperoxidase

AUTHOR(S): Yang, J. J.; Jennette, J. C.; Falk, R. J.

CORPORATE SOURCE: Department Medicine, University of North Carolina,

Chapel Hill, NC, 27599-7155, USA

SOURCE: Clinical and Experimental Immunology (1994),

97(3), 466-73

CODEN: CEXIAL; ISSN: 0009-9104

DOCUMENT TYPE: Journal LANGUAGE: English

Anti-neutrophil cytoplasmic antibodies (ANCA), including anti-myeloperoxidase (MPO) antibodies, are associated with pauci-immune necrotizing small vessel vasculitis or glomerulonephritis. To substantiate a pathogenic role for ANCA, an animal model of pauci-immune ANCA-induced glomerulonephritis or vasculitis is required. Brouwer et al. reported pauci-immune glomerulonephritis in rats immunized with human MPO followed by perfusion of kidneys with lysosomal enzyme extract combined with H2O2, and suggested that this could serve as a model of ANCA-induced disease. These studies were repeated here in spontaneously hypertensive rats (SHR) and Brown Norway rats (BNR). Rats were immunized with human When circulating anti-MPO antibodies were detectable by indirect immunofluorescence microscopy and ELISA, blood pressure was measured, then perfusion of the left kidney of each rat was done via the renal artery in a closed, blood-free circuit with either MPO + H2O2, MPO, H2O2 alone or MPO + H2O2 + neutral protease. Rats were killed on day 4 or day 10 after perfusion, and specimens were examined by light and immunofluorescence microscopy. Pathol. lesions and deposits of IgG, C3, and MPO were found in immunized rats perfused with MPO + H2O2 with or without neutral protease, or MPO alone, in both rat strains and on both day 4 and day 10. The degree of histol. injury was proportional in intensity to the amount of IgG immune deposits. Spontaneously hypertensive rats sustained more damage and higher blood pressure than Brown Norway rats. No lesion was observed in immunized rats perfused with H2O2 or in the non-perfused right kidneys. Some of the non-immunized rats perfused with MPO + H2O2 developed pathol. lesions. In conclusion, these rat models are examples of immune complex-mediated glomerulonephritis, and therefore are not similar to human ANCA-associated disease.

IT 9003-99-0, Myeloperoxidase RL: ADV (Adverse effect, including toxicity); BIOL (Biological study) RN 9003-99-0 CAPLUS

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

2843 REFERENCES IN FILE CA (1907 TO DATE)

132 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

2868 REFERENCES IN FILE CAPLUS (1907 TO DATE)

21 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

=> d his

L1

L3

(FILE 'HOME' ENTERED AT 10:56:25 ON 08 FEB 2007)

FILE 'REGISTRY' ENTERED AT 11:04:58 ON 08 FEB 2007

STRUCTURE UPLOADED

L2 7462 S L1 SSS FULL

FILE 'CAPLUS, USPATFULL' ENTERED AT 11:05:33 ON 08 FEB 2007

853 S L2 AND (HYPERTENSION OR HYPERTENSIVE OR HIGH BLOOD PRESSURE

L4 510 S L3 AND (HYPERTENSION OR HIGH BLOOD PRESSURE)

L5 611 S L3 AND (HYPERTENSION OR HIGH BLOOD PRESSURE OR ANTIHYPERTENSI

L6 586 DUP REM L5 (25 DUPLICATES REMOVED)

L7 571 S L6 NOT FERULIC ACID

L8 571 FOCUS L7 1-

L9 382 S L8 AND PD <= 2001

L10 382 FOCUS L9 1-

L11 378 S L10 NOT CAFFEIC ACID

L12 309 S L11 AND (HYPERTENSION OR HIGH BLOOD PRESSURE)

FILE 'REGISTRY' ENTERED AT 11:22:29 ON 08 FEB 2007

L13 45 S 3 CAFFEOYLQUINIC ACID OR NEOCHLORGENIC ACID 4 CAFFEOYLQUINIC

L14 65 S L13 OR DICAFFEOYL QUINIC ACID OR FERULOYLQUINIC ACID OR FERUL

L15 9 S DIMETHYL CAFFEATE ETHER OR PHENYLETHYL CAFFEATE OR CAFFEOYL A

L16 183 S DICAFFEOYLQUINIC ACID OR CICHORIC ACID OR CONIFERYL ALCOHOL O

L17 518066 S 14 OR L15 OR L16

L18 227 S L14 OR L15 OR L16

L19 66 S CURCUMIN

=> 118 not curcumin

L18 IS NOT A RECOGNIZED COMMAND

The previous command name entered was not recognized by the system. For a list of commands available to you in the current file, enter

"HELP COMMANDS" at an arrow prompt (=>).

=> s l18 not curcumin

66 CURCUMIN

L20 227 L18 NOT CURCUMIN

=> s l16 not curcumin

66 CURCUMIN

L21 183 L16 NOT CURCUMIN

=> file caplus

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST 321.15 1302.18

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL

CA SUBSCRIBER PRICE ENTRY SESSION 0.00 -92.04

FILE 'CAPLUS' ENTERED AT 11:29:37 ON 08 FEB 2007
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FILE COVERS 1907 - 8 Feb 2007 VOL 146 ISS 7 FILE LAST UPDATED: 7 Feb 2007 (20070207/ED)

Effective October 17, 2005, revised CAS Information Use Policies apply. They are available for your review at:

http://www.cas.org/infopolicy.html

=> s 118

L22 62987 L18

=> s 122 and (hypertension or high blood pressure or antihypertensives)

83810 HYPERTENSION

104 HYPERTENSIONS

83830 HYPERTENSION

(HYPERTENSION OR HYPERTENSIONS)

3972522 HIGH

591 HIGHS

3972854 HIGH

(HIGH OR HIGHS)

1287475 BLOOD

1248 BLOODS

1287616 BLOOD

(BLOOD OR BLOODS)

1226951 PRESSURE

176764 PRESSURES

1293696 PRESSURE

(PRESSURE OR PRESSURES)

2360 HIGH BLOOD PRESSURE

(HIGH (W) BLOOD (W) PRESSURE)

30815 ANTIHYPERTENSIVES

L23 159 L22 AND (HYPERTENSION OR HIGH BLOOD PRESSURE OR ANTIHYPERTENSIVE S)

=> focus

PROCESSING COMPLETED FOR L23

L24 159 FOCUS L23 1-

 \Rightarrow s 124 and pd \Rightarrow 001

L25 159 S L24

57 L25 AND PD <=2001

=> d ibib abs 1-57 hitstr

L26 ANSWER 1 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2002:90556 CAPLUS

DOCUMENT NUMBER: 136:131255

TITLE: Methods for early diagnosis of kidney disease and

treatment by drug intervention using lysosome

activating compounds

INVENTOR(S): Comper, Wayne D.

PATENT ASSIGNEE(S): Austria

SOURCE: U.S. Pat. Appl. Publ., 30 pp., Cont.-in-part of U.S.

Ser. No. 415,217. CODEN: USXXCO

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

| 1 | PATENT NO. | | | | | KIND DATE | | | APPLICATION NO. | | | | | | DATE | | | |
|-------|------------|------|------|------|-----|-----------|-----|------|-----------------|-----|-------|------|----------|--------|------|------|------|---------|
| τ | us Us | 2002 | 0129 | 06 | | A1 | - | 2002 | 0131 | | US 2 | 001- | 8933 | 46 | | 2 | 0010 | 628 |
| Ţ | US | 2002 | 1107 | 99 | | A1 | | 2002 | 0815 | | US 1 | 999- | 4152 | 17 | | 1: | 9991 | 012 |
| τ | US | 6447 | 989 | | | В2 | | 2002 | 0910 | | | | | | | | | |
| 7 | WO | 2000 | 0379 | 44 | | A1 | | 2000 | 0629 | | WO 1 | 999- | IB20 | 29 | | 1: | 9991 | 220 < |
| | | W: | ΑE, | AL, | AM, | AT, | ΑU, | AZ, | BA, | BB, | ВG, | BR, | BY, | CA, | CH, | CN, | CR, | CU, |
| | | | | | | | | ES, | | | | | | | | | | |
| | | • | | | | | | KP, | | | | | | | | | | |
| | | | MG, | MK, | MN, | MW, | MX, | NO, | NZ, | PL, | PT, | RO, | RU, | SD, | SE, | SG, | SI, | SK, |
| | | | SL, | ТJ, | TM, | TR, | TT, | TZ, | UA, | UG, | UΖ, | VN, | ΥU, | ZA, | ZW | | | |
| | | RW: | GH, | GM, | KE, | LS, | MW, | SD, | SL, | SZ, | TZ, | ŬĠ, | ZW, | AT, | BE, | CH, | CY, | DE, |
| | | | DK, | ES, | FI, | FR, | GB, | GR, | ΙE, | IT, | LU, | MC, | NL, | PT, | SE, | BF, | ВJ, | CF, |
| | | | CG, | CI, | CM, | GΑ, | GN, | GW, | ML, | MR, | ΝE, | SN, | TD, | TG | | | | |
| 2 | ZA | 2001 | 0050 | 58 | | Α | | 2002 | 0620 | | ZA 2 | 001- | 5058 | | | 2 | 0010 | 620 |
| Ţ | US | 2004 | 1061 | 55 | | A1 | | 2004 | 0603 | • | US 2 | 003- | 7213 | 51 | | 2 | 0031 | 126 |
| Č | JP | 2006 | 0388 | 77 | | Α | | 2006 | 0209 | 1 | JP 2 | 005- | 2701 | 60 | | 2 | 0050 | 916 |
| PRIOR | ITY | APP | LN. | INFO | . : | | | | | | AU 1 | 998- | 7843 | | | A 19 | 9981 | 221 |
| | | | | | | | | | | | US 1 | 999- | 4152 | 17 | | A2 1 | 9991 | 012 |
| | | | | | | | | | | • | WO 1. | 999- | IB20: | 29 | 1 | W 19 | 9991 | 220 |
| | | | | | | | | | | | JP 2 | 000- | 5899 | 50 | | A3 1 | 9991 | 220 |
| | | | | | | | | | | • | US 2 | 001- | 8933 | 46 | | A1 2 | 0010 | 628 |

AB A method is disclosed for diagnosing early stage of a disease in which an intact protein found in urine is an indicator of the disease, followed by early drug intervention to prevent and treat the disease are also disclosed. The drug treatment involves the use of a lysosome activating compound Urine samples of normal and diabetic patients were analyzed by size-exclusion chromatog. and HPLC.

IT 9003-99-0, Peroxidase

RL: ANT (Analyte); DGN (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)

(horseradish; methods for early diagnosis of kidney disease and treatment by drug intervention using lysosome activating compds.)

RN 9003-99-0 CAPLUS

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***.

L26 ANSWER 2 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2001:903829 CAPLUS

DOCUMENT NUMBER: 136:15240

TITLE: Method for treating hyperglycemia and conditions

associated with damage caused by reducing sugars using

aminoquanidine

Wuerth, Jean-Paul; Cartwright, Kenneth INVENTOR(S):

PATENT ASSIGNEE(S): SOURCE:

Alteon, Inc., USA PCT Int. Appl., 20 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| I | PATE | NT 1 | 10. | | | KINI |) | DATE | | 1 | APPL | ICAT: | ION 1 | . O <i>l</i> | | Di | ATE | |
|--------|-------|------|-------|-----|-----|------|-----|------|------|-----|-------|-------|-------|--------------|-----|-------|-------|-------|
| - | | | | | | | - | | , - | | | | | | | - | | |
| | WO 2 | 0010 | 9389 | 54 | | A1 | | 2001 | 1213 | 1 | WO 2 | 001- | US408 | 374 | | 20 | 0010 | 507 < |
| | Ţ | W: | ΑE, | AG, | AL, | AM, | ΑT, | AU, | ΑZ, | BA, | BB, | BG, | BR, | BY, | ΒZ, | CA, | CH, | CN, |
| | | | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EE, | ES, | FI, | GB, | GD, | GE, | GH, | GM, |
| | | | HR, | ΗU, | ID, | IL, | IN, | IS, | JΡ, | KΕ, | KG, | ΚP, | KR, | ΚZ, | LC, | LK, | LR, | LS, |
| | | | LT, | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | MZ, | NO, | ΝZ, | PL, | PT, | RO, |
| | | | RU, | SD, | SE, | SG, | SI, | SK, | SL, | ТJ, | TM, | TR, | TT, | ΤZ, | UA, | ŪĠ, | UZ, | VN, |
| | | | YU, | ZA, | ZW, | AM, | ΑZ, | BY, | KG, | ΚZ, | MD, | RU, | ТJ, | TM | | | | |
| | J | RW: | GH, | GM, | KE, | LS, | MW, | ΜZ, | SD, | SL, | SZ, | TZ, | UG, | ZW, | ΑT, | BE, | CH, | CY, |
| | | | DE, | DK, | ES, | FI, | FR, | GB, | GR, | ΙE, | ΙT, | LU, | MC, | NL, | PT, | SE, | TR, | BF, |
| | | | ВJ, | CF, | CG, | CI, | CM, | GA, | GN, | GW, | ML, | MR, | ΝE, | SN, | TD, | TG | | |
| Ţ | JS 20 | 0030 | 4558 | 32 | | A1 | | 2003 | 0306 | 1 | US 20 | 002- | 7185 | 1 | | 20 | 00202 | 208 |
| PRIORI | ITY A | APPI | JN.] | NFO | . : | | | | | 1 | US 20 | 000- | 21013 | 14P |] | P 20 | 0000 | 507 |
| | | | | | | | | | | 1 | US 20 | 001- | 87687 | 74 | 1 | 31 20 | 00106 | 507 |

Provided, among other things, are methods for treating mammals, such as AB humans, with diabetes mellitus to delay the onset of end stage renal disease, relating to administering an effective amount of a pharmaceutical composition, wherein said composition comprise, a compound selected from the group

consisting of aminoguanidine, its pharmaceutically acceptable salts, and mixts. thereof. Methods are disclosed for the gradual administration of the compound for treatment of diabetes or other indications associated with damage caused by reducing sugars, for the use of a periodic screening test for crescentic glomerulonephritis, and for treating humans with indicia of overt diabetic nephropathy.

IT 9003-99-0, Myeloperoxidase

> RL: BSU (Biological study, unclassified); BIOL (Biological study) (anti-neutrophil cytoplasmic antibodies of the myeloperoxidase-type; aminoguanidine treatment to delay the onset of end stage renal disease in diabetic subjects screened for crescentic glomerulonephritis by measuring MPO-ANCA levels)

RN9003-99-0 CAPLUS

SOURCE:

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 3 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2001:902404 CAPLUS

DOCUMENT NUMBER: 136:163044

TITLE: Interactions of nitric oxide-derived reactive nitrogen

species with peroxidases and lipoxygenases

AUTHOR (S): Coffey, Marcus J.; Coles, Barbara; O'Donnell, Valerie

CORPORATE SOURCE: Wales Heart Research Institute, University of Wales

College of Medicine, Cardiff, CF14 4XN, UK Free Radical Research (2001), 35(5), 447-464

CODEN: FRARER; ISSN: 1071-5762

PUBLISHER: Harwood Academic Publishers DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

A review. Nitric oxide (NO) is a major free radical modulator of smooth AB muscle tone, which under basal conditions acts to preserve vascular homeostasis through its anti-inflammatory properties. The biochem. of NO, in particular, its rapid conversion in vivo into secondary reactive nitrogen species (RNS), its chemical nature as a free radical and its high diffusibility and hydrophobicity dictate that this species will interact with numerous biomols. and enzymes. In this review, the authors consider the interactions of a number of enzymes found in the vasculature with NO and NO-derived RNS. All these enzymes are either homeostatic or promote the development of atherosclerosis and hypertension. Therefore their interactions with NO and NO-derived RNS will be of central importance in the initiation and progression of vascular disease. In some examples, (e.g., lipoxygenase, LOX), such interactions provide catalytic "sinks" for NO, but for others, in particular peroxidases and prostaglandin H synthase (PGHS), reactions with NO may be detrimental. Nitric oxide and NO-derived RNS directly modulate the activity of vascular peroxidases and LOXs through a combination of effects, including transcriptional regulation, altering substrate availability, and direct reaction with enzyme turnover intermediates. Therefore, these interactions will have two major consequences: (i) depletion of NO levels available to cause vasorelaxation and prevent leukocyte/platelet adhesion and (ii) modulation of activity of the target enzymes, thereby altering the generation of bioactive signaling mols. involved in maintenance of vascular homeostasis, including prostaglandins and leukotrienes.

IT 9003-99-0, Peroxidase

RL: BSU (Biological study, unclassified); BIOL (Biological study) (interactions of nitric oxide-derived reactive nitrogen species with peroxidases and lipoxygenases)

RN 9003-99-0 CAPLUS

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT:

174 THERE ARE 174 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 4 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2001:826669 CAPLUS

DOCUMENT NUMBER: 137:57317

TITLE: Effect of intravenous taurine on endotoxin-induced

acute lung injury in sheep

AUTHOR(S): Egan, Bridget M.; Abdih, Hazem; Kelly, Cathal J.;

Condron, Claire; Bouchier-Hayes, David J.

Condron, Claire; Bouchier-Hayes, David J.

CORPORATE SOURCE: Department of Surgery, Beaumont Hospital, Dublin, Ire.

SOURCE: European Journal of Surgery (2001), 167(8),

575-580

CODEN: EUJSEH; ISSN: 1102-4151

PUBLISHER: Taylor & Francis Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

Objective: To find out if pretreatment with taurine would reduce the severity of endotoxin-induced acute lung injury in a large animal model. Design: Randomized controlled study under license from the Department of Health. Setting: Department of Surgical Research, Ireland. Animals: 15 male Suffolk sheep. Interventions: Vascular catheters were placed in the femoral artery and vein and a Swan-Ganz catheter in the external jugular vein under general anesthetic. Animals were randomized into 3 groups: control with measurements taken at baseline and half hourly up to 90 min; endotoxin, given Escherichia coli endotoxin i.v. after baseline measurements and taurine given 300 mg/kg 1 h before endotoxin was given. Main outcome measures: Mean systemic arterial pressure, mean pulmonary arterial pressure, arterial oxygen tension (pO2), pulmonary myeloperoxidase activity, and neutrophil respiratory burst activity. Results: Endotoxin induced a severe lung injury characterized by a

decrease in mean systemic blood pressure and an increase in pulmonary artery pressure, hypoxia, and an increase in pulmonary myeloperoxidase activity. Pretreatment with i.v. taurine significantly reduced these hemodynamic changes. It reduced pulmonary myeloperoxidase activity and peripheral neutropenia and increased neutrophil respiratory burst activity. Conclusions: This data suggest that taurine may have a therapeutic role in preventing the lung injury seen in endotoxemia. 9003-99-0, Myeloperoxidase

RL: BSU (Biological study, unclassified); BIOL (Biological study) (effect of i.v. taurine on pulmonary myeloperoxidase activity after endotoxin-induced acute lung injury in sheep)

RN 9003-99-0 CAPLUS

IT

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT: 23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 5 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2001:816459 CAPLUS

DOCUMENT NUMBER: 135:339302

TITLE: Methods and compositions for enhancing cellular

function through protection of tissue components
INVENTOR(S): Frey, William H., II; Fawcett, John Randall; Thorne,

Robert Gary; Chen, Xueqing

PATENT ASSIGNEE(S): Healthpartners Research Foundation, USA

SOURCE: PCT Int. Appl., 77 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| | PATENT NO. | | | | APPLICATION NO. | | | | | DATE | | | | | | | |
|----------|--------------------------|---------------------------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|--|
| WO | 20010829 | 32 | | A2 | | 2001 | 1108 | 1 | | | | | | 2 | 0010 | 430 < | |
| | CR, HU, LU, SD, | AG, CU, ID, LV, SE, ZW | CZ, IL, MA, | DE, IN, MD, | DK, IS, MG, | DM, JP, MK, | DZ, KE, MN, | EE, KG, MW, | ES, KP, MX, | FI, KR, MZ, | GB, KZ, NO, | GD, LC, NZ, | GE, LK, PL, | GH, LR, PT, | GM, LS, RO, | HR, LT, RU, | |
| IIS. | • | DK, CF, | ES, CG, | FI, CI, | FR, CM, | GB, GA, | GR, GN, | IE, GW, | IT, ML, | LU, MR, | MC, NE, | NL, SN, | PT, TD, | SE, TG | TR, | BF, | |
| US | 7084126 | 00 | | B2 | | 2002 | 0307 | ! | 05 2 | 001- | 0444. | 50 | | 4 | 0010 | 12/ | |
| CA | 2429162 | | | A1 | | 2001 | 1108 | . (| CA 2 | 001- | 2429 | 162 | | 2 | 0010 | 430 < | |
| EP | 2429162 1278525 | • | | A2 | • | 2003 | 0129 |] | EP 2 | 001- | 9309 | 57 | | 2 | 0010 | 130 | |
| | 1278525 | | | | | | | | | | | | | | | | |
| | R: AT, IE, | BE, SI, | | | | | | | | | LI, | LU, | ΝL, | SE, | MC, | PT, | |
| | 344040 | | | | | | | | | | | | | | | | |
| US | 20052726 | 42 | | A1. | : | 2005 | 1208 | 1 | US 2 | 005- | 1919 | 01 | | 2 | 0050 | 728 | |
| | 20060094 | | | | | | | | | | | | | | | | |
| | 20060094 | | | | | | | | | | | | | | | | |
| US | 20060147 | 16 | | Al | | 2006 | 0119 |] | JS 2 | 005- | 2202 | 23 | | 2 | 0050 | 906 | |
| | 20060305 | | | ΑI | | 2006 | 0209 | | | | | | | | | | |
| PRIORITI | APPLN. | INFO | . : | | | | | | | 000- | | | | | | | |
| | | | | | | | | | | 000- | | | | | | | |
| | | | | | | | | Ţ | JS 2 | 001-1 001-1 | 8444 | 50 | i | A3 2 | 0010 | 127 | |

OTHER SOURCE(S): MARPAT 135:339302

AB Methods and compns. for enhancing cellular function through protection of tissue components, such as receptors, proteins, lipids, nucleic acids, carbohydrates, hormones, vitamins, and cofactors, by administering pyrophosphate analogs or related compds. Preferably, the invention provides a method for protecting a muscarinic acetylcholine receptor (mAChR) an/or increasing the efficacy of and agent the directly or indirectly affects a mAChR in a subject in need thereof.

IT 9003-99-0, Peroxidase

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(methods and compns. for enhancing cellular function through protection of tissue components such as muscarinic receptors by administering pyrophosphate analogs and combination with other agents)

RN 9003-99-0 CAPLUS

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

L26 ANSWER 6 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2001:338762 CAPLUS

DOCUMENT NUMBER: 134:362292

TITLE: Methods of determining individual hypersensitivity to

a pharmaceutical agent from gene expression profile

INVENTOR(S): Farr, Spencer

PATENT ASSIGNEE(S): Phase-1 Molecular Toxicology, USA

SOURCE: PCT Int. Appl., 222 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PAT | CENT : | NO. | | | KIN | D : | DATE | | | APPL | ICAT | ION : | NO. | | D | ATE | |
|----------|--------|----------|--------|-----|-----|-----|----------|----------|-----|----------|-----------|-----------|---------|-----|-----|----------|-----------|
| WO. | 2001 | 0329 | 28 | | A2 | _ | 2001 | 0510 | , | WO 2 | 000-1 | US30: | 474 | | 2 | 0001 | 103 < |
| | 2001 | | | | A3 | | 2002 | | | | | | - / - | | _ | | |
| | W : | ΑE, | AG, | AL, | AM, | ΑT, | ΑU, | AZ, | BA, | BB, | BG, | BR, | BY, | ΒZ, | CA, | CH, | CN, |
| | | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EE, | ES, | FI, | GB, | GD, | GE, | GH, | GM, | HR, |
| | | HU, | ID, | IL, | IN, | IS, | JP, | KE, | KG, | ΚP, | KR, | KΖ, | LC, | LK, | LR, | LS, | LT, |
| | | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | MZ, | NO, | NZ, | PL, | PT, | RO, | RU, |
| | | SD, | SE, | SG, | SI, | SK, | SL, | ТJ, | TM, | TR, | TT, | TZ, | UA, | ŪĠ, | US, | UZ, | VN, |
| | | ΥU, | ZA, | ZW, | AM, | ΑZ, | BY, | KG, | ΚZ, | MD, | RU, | TJ, | TM | | | | |
| | RW: | GH, | GM, | ΚE, | LS, | MW, | MZ, | SD, | SL, | SZ, | TZ, | UG, | ZW, | ΑT, | BE, | CH, | CY, |
| | | | | | | | GB, | | | | | | | | | | |
| | | ВJ, | CF, | CG, | CI, | CM, | GA, | GN, | GW, | ML, | MR, | NE, | SN, | TD, | TG | - | • |
| PRIORITY | APP | | | | | | | · | | | | | | | P 1 | 9991 | 105 |
| | | | | | | | | | 1 | JS 2 | 000- | 1965 | 71P |] | P 2 | 0000 | 111 |

The invention discloses methods, gene databases, gene arrays, protein AB arrays, and devices that may be used to determine the hypersensitivity of individuals to a given agent, such as drug or other chemical, in order to prevent toxic side effects. In one embodiment, methods of identifying hypersensitivity in a subject by obtaining a gene expression profile of multiple genes associated with hypersensitivity of the subject suspected to be hypersensitive, and identifying in the gene expression profile of the subject a pattern of gene expression of the genes associated with hypersensitivity are disclosed. The gene expression profile of the subject may be compared with the gene expression profile of a normal individual and a hypersensitive individual. The gene expression profile of the subject that is obtained may comprise a profile of levels of mRNA or cDNA. The gene expression profile may be obtained by using an array of nucleic acid probes for the plurality of genes associated with hypersensitivity. The expression of the genes predetd. to be associated with hypersensitivity is directly related to prevention or repair of toxic damage at the tissue, organ or system level. Gene databases arrays and apparatus useful for identifying hypersensitivity in a subject are also disclosed.

9003-99-0, Myeloperoxidase TT

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(methods of determining individual hypersensitivity to a pharmaceutical

agent

from gene expression profile)

9003-99-0 CAPLUS RN

Peroxidase (9CI) (CA INDEX NAME) CN

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

L26 ANSWER 7 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2001:247215 CAPLUS

DOCUMENT NUMBER: 134:276498

Engineering of replication selective adenoviruses with TITLE:

tumor-associated antigen promoter for use in cancer

APPLICATION NO. DATE

therapy

Molnar-kimber, Katherine; Toyoizumi, Takane INVENTOR(S):

PATENT ASSIGNEE(S): The Trustees of the University of Pennsylvania, USA

DATE

SOURCE: PCT Int. Appl., 56 pp.

direct, oncolytic effect on the tumor.

CODEN: PIXXD2

DOCUMENT TYPE:

LANGUAGE:

IT

Patent English

KIND

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.

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A1 20010405 WO 2000-US27212 20001002 <--
     WO 2001023004
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
             HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
             LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
            SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
             YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
             CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.:
                                           US 1999-157224P
                                                             P 19990930
    The invention provides a replication selective adenovirus (Ad) mutant with
     improved selectively for tumor cells expressing the tumor associated antigen
     in cancers and malignancies, as well as in proliferative cells,
     characterizing diseases, such as restenosis, intimal proliferative disease
     and pulmonary hypertension. The selected Ad vectors are driven
    by promoters of the tumor associated antigens, or RNA transcripts or genes
    therefor, substituting for the activity of at least adenovirus E1A
    promoter, which has been deactivated or diminished. Also provided is the
    use of the Ad vector to deliver therapeutic compns. to patients, as well
    as a method for treating cancers, such as CEA pos. cancers, or
    proliferative cell diseases in a patient by administering to the patient
    an effective amount of the Ad vector, which may also express a therapeutic
    gene or peptide, and treatment may also be combined with radiation,
    chemotherapy or immunomodulatory agents. The Ad is designed to replicate
    within the tumor cell, thereby spreading throughout the tumor nodule.
    This permits the delivery of a much higher dose of the heterologous
    therapeutic protein than previously possible, and the virus achieves a
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9002-10-2, Tyrosinase RL: BSU (Biological study, unclassified); BIOL (Biological study)

(promoter, specific to tumor expressing; engineering of replication

selective adenoviruses with tumor-associated antigen promoter for use in cancer therapy)

RN 9002-10-2 CAPLUS

CN Oxygenase, monophenol mono- (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 8 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2001:44267 CAPLUS

DOCUMENT NUMBER: 134:290172

TITLE: Effect of captopril on mushroom tyrosinase activity in

vitro

AUTHOR(S): Espin, J. C.; Wichers, H. J.

CORPORATE SOURCE: Laboratorio de Fitoquimica, Departamento de Ciencia y

Tecnologia de Alimentos, CEBAS-CSIC, Murcia, 30080,

Spain

SOURCE: Biochimica et Biophysica Acta, Protein Structure and

Molecular Enzymology (2001), 1544(1-2),

289-300

CODEN: BBAEDZ; ISSN: 0167-4838

PUBLISHER: Elsevier B.V.

DOCUMENT TYPE: Journal LANGUAGE: English

The study presented here demonstrates that the antihypertensive drug captopril ([2S]-N-[3-mercapto-2-methylpropionyl]-L-proline) is an irreversible non-competitive inhibitor and an irreversible competitive inhibitor of the monophenolase and diphenolase activities of mushroom tyrosinase when L-tyrosine and L-DOPA were assayed spectrophotometrically in vitro, resp. Captopril was rendered unstable by tyrosinase catalysis because of the interaction between the enzymic-generated product (o-quinone) and captopril to give rise to a colorless conjugate. Therefore, captopril was able to prevent melanin formation. The spectrophotometric recordings of the inhibition of tyrosinase by captopril were characterized by the presence of a lag period prior to the attainment of an inhibited steady state rate. The lag period corresponded to the time in which captopril was reacting with the enzymically generated o-quinone. Increasing captopril concns. provoked longer lag periods as well as a concomitant decrease in the tyrosinase activity. Both lag period and steady state rate were dependent of captopril, substrate and tyrosinase concns. The inhibition of both monophenolase and diphenolase activities of tyrosinase by captopril showed pos. kinetic co-operativity which arose from the protection of both substrate and o-quinone against inhibition by captopril. Inhibition expts. carried out using a latent mushroom tyrosinase demonstrated that captopril only bound the enzyme at its active site. The presence of Cu ions only partially prevented but not reverted mushroom tyrosinase inhibition. This could be due to the formation of both Cu-captopril complex and disulfide interchange reactions between captopril and cysteine rich domains at the active site of the enzyme.

IT 9002-10-2

RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(effect of captopril on mushroom tyrosinase activity in vitro)

RN 9002-10-2 CAPLUS

CN Oxygenase, monophenol mono- (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT: 49 THERE ARE 49 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 9 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2000:573694 CAPLUS

DOCUMENT NUMBER:

133:182988

TITLE:

Organosilicate sol-gel matrixes for drug delivery

INVENTOR(S):

Babich, John W.; Bonavia, Grant; Zubieta, Jon

PATENT ASSIGNEE(S):

Biostream, Inc., USA

SOURCE:

PCT Int. Appl., 133 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

Patent English

DANGUAGE.

UNT: 1

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE _____ _____ ______ _____ _ _ _ _ 20000817 WO 2000-US3754 WO 2000047236 A1 20000214 <--W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG CA 2328614 Α1 20000817 CA 2000-2328614 20000214 <--US 6395299 В1 20020528 US 2000-503438 20000214 Т JP 2002536422 20021029 JP 2000-598187 20000214 AU 772153 B2 20040408 AU 2000-27599 20000214 US 2003082238 A1 20030501 US 2002-77475 20020215 US 2004241205 A1 20041202 US 2004-838423 20040504 US 7052913 B2 20060530 PRIORITY APPLN. INFO.: US 1999-119828P P 19990212 US 2000-503438 A1 20000214 WO 2000-US3754 W 20000214

US 2002-77475 Al 20020215

AB Biocompatible matrixes such as sol-gels encapsulating a reaction center may be administered to a subject for conversion of prodrugs into biol. active agents. In certain embodiments, the biocompatible matrixes of the present invention are sol-gels. In one embodiment, the enzyme L-amino acid decarboxylase is encapsulated and implanted in the brain to convert L-dopa to dopamine for treatment of Parkinson's disease. The silica sol was prepared by the addition of substituted trimethoxysilanes, tetra-Me orthosilicate (TMOS) and 4 mM HCl solution Total desired volume of the sol was determined by the number of matrixes to be prepared Entrapment of penicillinase in

the matrix was performed by using pH 6.5 phosphate buffer. The penicillinase activity was determined by using 3 mM solution of penicillin G in buffer.

IT 9002-10-2, Tyrosinase

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (organosilicate sol-gel matrixes for drug delivery)

RN 9002-10-2 CAPLUS

CN Oxygenase, monophenol mono- (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT:

8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 10 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2000:534984 CAPLUS

DOCUMENT NUMBER:

133:144942

TITLE:

Melanogenesis-stimulating mono- and bicyclic monoterpene diols as dermatological compounds,

preparation, and use

INVENTOR(S): Ren, Wu Yun; Brown, David A.

PATENT ASSIGNEE(S): Codon Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 40 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

3

LANGUAGE:
FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

KIND DATE APPLICATION NO. DATE PATENT NO. _____ --------------------WO 1999-US11841 19990528 <--20000803 WO 2000044368 **A1** W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG US 6214888 В1 20010410 US 1998-86547 19980528 <--AU 9942155 20000818 A1 AU 1999-42155 19990528 <--PRIORITY APPLN. INFO.: US 1998-86547 A 19980528 US 1996-26577P P 19960918 US 1997-35947P P 19970121 US 1997-36863P P 19970204 US 1997-48597P P 19970604 WO 1997-US16642 A2 19970918 A1 19980318 WO 1998-US5346 WO 1999-US11841 W 19990528

OTHER SOURCE(S): MARPAT 133:144942

AB Monocyclic and bicyclic monoterpene diols are provided that stimulate melanogenesis in mammalian skin, hair, wool or fur, and are useful for treating or preventing various skin and proliferative disorders, neurodegenerative diseases, and diseases regulated by the nitric oxide/cyclic GMP/protein kinase G pathway.

IT 9002-10-2, Tyrosinase

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(melanogenesis-stimulating mono- and bicyclic monoterpene diols as dermatol. compds., preparation, and use)

RN 9002-10-2 CAPLUS

CN Oxygenase, monophenol mono- (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT:

4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 11 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2000:442020 CAPLUS

DOCUMENT NUMBER:

133:57150

TITLE:

The diagnosis and monitoring of treatment for the

early stages of renal disease and/or renal

complications of disease through the determination of proteinuria using immunological or non-immunological

techniques

INVENTOR(S):

Comper, Wayne D.

PATENT ASSIGNEE(S):

Monash University, Australia

SOURCE:

PCT Int. Appl., 50 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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PATENT NO.
                        KIND
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                                        WO 1999-IB2029
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            MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK,
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        RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
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            CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                         US 1999-415217
    US 2002110799
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                                                                19991012
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    CA 2356174
                               20000629
                                                                19991220 <--
                                          BR 1999-16407
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    EP 1141728
                         Α1
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            IE, SI, LT, LV, FI, RO
                                          JP 2000-589950
    JP 2002533680
                        т
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                                                                19991220
                                         ZA 2001-5058
    ZA 2001005058
                         Α
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                                                                20010620
                        A1
    US 2002012906
                              20020131
                                          US 2001-893346
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                        A1
    US 2004106155
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                                          US 2003-721351
                                                                20031126
    JP 2006038877
                        Α
                               20060209
                                          JP 2005-270160
                                                                20050916
PRIORITY APPLN. INFO.:
                                          AU 1998-7843
                                                             A 19981221
                                          US 1999-415217
                                                            A 19991012
                                          JP 2000-589950
                                                            A3 19991220
                                          WO 1999-IB2029
                                                            W 19991220
                                          US 2001-893346
                                                            A1 20010628
AB
```

AB A method is disclosed for diagnosing early stage of a disease in which an intact protein found in urine is an indicator of the disease. The method includes assaying urine sample to detect the presence of modified protein using either immunol. or non-immunol. technique. Methods for preventing and treating the disease are also disclosed.

IT 9003-99-0, Peroxidase

RL: ANT (Analyte); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)

(horseradish; diagnosis and monitoring of treatment for early stages of renal disease and/or renal complications of disease through determination of proteinuria using immunol. or non-immunol. techniques)

RN 9003-99-0 CAPLUS

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 12 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2000:277810 CAPLUS

DOCUMENT NUMBER:

132:326056

TITLE: INVENTOR(S): Systems for oral delivery Russell-Jones, Gregory John

PATENT ASSIGNEE(S):

Biotech Australia Pty. Ltd., Australia

SOURCE:

PCT Int. Appl., 32 pp.

DOCUMENT TYPE:

CODEN: PIXXD2

LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---------------|------|----------|-----------------|------------|
| | | | | |
| WO 2000022909 | A2 | 20000427 | WO 1999-IB1872 | 19991018 < |

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WO 2000022909
                           A3
                                 20001123
            AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,
             CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL,
             IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA,
             MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ,
             BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
             DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
             CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     AU 2000010712
                                 20000508
                                             AU 2000-10712
                          Α
                                                                     19991018 <--
PRIORITY APPLN. INFO.:
                                             US 1998-104827P
                                                                  P 19981019
                                             WO 1999-IB1872
                                                                  W 19991018
     A pharmaceutical and a biol. active substance, for oral administration,
AB
     can be "coated" or "encapsulated" with a carboxylic acid, such that the
     substance is protected from proteolysis in the stomach and is taken up
     from the intestine. It is thought that the carboxylic acids coat and
     protect the active agent from the proteolytic environment of the stomach,
     allowing the agent to pass safely through the stomach and to be absorbed
     in the small intestines. The carboxylic acid agent complex can be adopted
     for oral, nasal, buccal, and transdermal delivery of moderately soluble and
     even insol. bioactive agents.
IT
     9003-99-0, Myeloperoxidase
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (carboxylic acids for encapsulating or enteric coating biol. active
        agents for delivery to intestine)
RN
     9003-99-0 CAPLUS
CN
     Peroxidase (9CI)
                       (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L26 ANSWER 13 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER:
                         2000:13671 CAPLUS
DOCUMENT NUMBER:
                         133:15695
TITLE:
                         Inhibition of polyadenosine diphosphate-ribose
                         synthetase attenuates dysfunction of pulmonary
                         vasorelaxation in acute lung injury
AUTHOR (S):
                         Pulido, Edward J.; Bensard, Denis D.; Shames, Brian
                         D.; Selzman, Craig H.; McIntyre, Robert C., Jr.
CORPORATE SOURCE:
                         Department of Surgery, University of Colorado Health
                         Sciences Center, Denver, CO, USA
SOURCE:
                         Surgical Forum (1998), 49, 14-15
                         CODEN: SUFOAX; ISSN: 0071-8041
PUBLISHER:
                         American College of Surgeons
DOCUMENT TYPE:
                         Journal
LANGUAGE:
                         English
     Acute respiratory distress syndrome (ARDS) is characterized by hypoxemia,
     decreased compliance, and pulmonary hypertension. The main
     purpose of this study was to determine the effect of polyadenosine
     diphosphate-ribose synthetase (PARS) inhibition by endotoxin (ETX) on
     pulmonary vasorelaxation and to determine the effect of PARS inhibition on
     neutrophil accumulation and edema in the lung. ETX-induced acute lung
     injury results in polymorphonuclear neutrophil-mediated dysfunction of
     pulmonary vasorelaxation. these data support the hypothesis that
     ETX-induced vascular dysfunction is mediated by cellular energy depletion
     as a result of excessive PARS activity.
ΙT
     9003-99-0, Myeloperoxidase
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (endotoxin-induced polyadenosine diphosphate-ribose synthetase
        inhibition and its relation to acute respiratory distress syndrome in
        human)
RN
     9003-99-0 CAPLUS
CN
     Peroxidase (9CI)
                      (CA INDEX NAME)
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*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 14 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 1999:600732 CAPLUS

DOCUMENT NUMBER: 132:117292

TITLE: The therapeutic effects of nitric oxide synthase

inhibitors, sulfasalazine or the proteasome inhibitor on the chronic intestinal and colonic inflammation

developed in HLA-B27 transgenic rats

AUTHOR(S): Aiko, Satoshi; Grisham, Matthew B.

CORPORATE SOURCE: Department of Surgery II, National Defense Medical

College, Louisiana State University Medical Center,

USA

SOURCE: Furi Rajikaru no Rinsho (1998), 13, 64-69

CODEN: FRRIFI

PUBLISHER: Nihon Igakukan

DOCUMENT TYPE: Journal LANGUAGE: Japanese

AB To determine the therapeutic effects of certain nitric oxide synthase (NOS) inhibitors, sulfasalazine (SZ) or the proteasome inhibitor, HLA-B27 male rats (B27 rats) that spontaneously developed colitis were randomized into six groups consisting of one untreated group, one vehicle group and four

treated groups that received either aminoguanidine (AG),

NG-nitro-L-arginine Me ester (L-NAME) or SZ in their drinking water, and MG-341, the selective proteasome inhibitor, orally for 21 days. Fisher 344 male rats were used as healthy control. We found that only AG was clin. effective on the colitic symptoms. Treatment with L-NAME resulted in deterioration of colitic symptoms, severe hypertension and enhanced mucosal permeabilities. Treatment with AG and SZ but not MG-341 attenuated the increases in ileal and colonic mucosal permeabilities. AG, L-NAME and SZ significantly attenuated the increases in the MPO activity in the distal colon, while MG-341 attenuated the MPO activity only in the proximal colon. We found that both AG and L-NAME but not SZ nor MG-341 significantly attenuated the increased plasma NO2-/NO3- levels. These results suggested that (1) the selective inhibition of iNOS may be more proper in order to manage the intestinal inflammation in the B27 rats, and (2) the granulocyte recruitment into the distal colon in this model may be

(2) the granulocyte recruitment into the distal colon in this model may be regulated by NO-dependent and -independent pathways. Addnl. studies will be required to determine the therapeutic effects of proteasome inhibitor in this model of chronic colitis.

IT 9003-99-0, Myeloperoxidase

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(response; therapeutic effects of nitric oxide synthase inhibitors, sulfasalazine or the proteasome inhibitor MG-341 on chronic intestinal and colonic inflammation developed in HLA-B27 transgenic rats)

RN 9003-99-0 CAPLUS

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

L26 ANSWER 15 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1999:595348 CAPLUS

DOCUMENT NUMBER: 131:225828

TITLE: Methods of diagnosis and triage using cell activation

measures

INVENTOR(S): Stoughton, Roland B.; Schmid-Schonbein, Geert W.;

Hugli, Tony E.; Kistler, Erik

PATENT ASSIGNEE(S): Cell Activation, Inc., USA; The Regents of the

University of California; The Scripps Research

Institute

SOURCE: PCT Int. Appl., 184 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| | PATENT NO. | | | | | | | APPLICATION NO. | | | | | | | | | |
|---------|--------------|-------------------|-------------------|-------------------|-------------------|-------------------|--------------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|
| WO | | 367 | | | A2 | | | | 1 | | | | | | | 9990 | 311 < |
| | W: | DE, JP, MW, | DK, KE, MX, | EE, KG, NO, | ES, KP, NZ, | FI, KR, PL, | AZ, GB, KZ, PT, | GD, LC, RO, | GE, LR, RU, | GH, LS, SD, | GM, LT, SE, | HR, LU, SG, | HU, LV, SI, | ID, MD, SK, | IL, MG, SL, | IN, MK, TJ, | IS, MN, TM, |
| | RW: | TJ, GH, ES, | TM GM, FI, | KE, | LS, GB, | MW, | UZ, SD, IE, | SL, | SZ, LU, | UG, | ZW, | AT, | BE, | CH, | CY, | DE, | DK, |
| | 2003 | 1903 | 68 | | A1 | | | 1009 | 1 | US 1 | 998- | 3889 | | | | | 311 311 < |
| UA | 9931 1062 | 829 323 | | | A A2 | | 1999 2000 | 0927 1227 | <i>i</i> 1 | AU 1: EP 1: | 999- 999- | 3182 9138 | 9 43 | | 1: 1: | 9990: 9990: | 311 < 311 < |
| | R: 2002 | IE, | si, | LT, | LV, | FI, | | • | · | • | · | · | · | · | • | • | , |
| PRIORIT | | | | | | | 2002 | | 1 | US 1: | 998- | 3889 US52 | 4 | 7 | A2 1 | 9980 | 311 |

AB Diagnostic methods that rely on the use of one or more assays that assess cellular activation are provided. The assays are performed on whole blood or leukocytes (neutrophils), and indicate individually or in combination the level of cardiovascular cell activation, which is pivotal in many chronic and acute disease states. These results of the assays are used within a clin. framework to support therapeutic decisions such as: further testing for infectious agents, anti-oxidant or anti-adhesion therapy, postponement and optimal re-scheduling of high-risk surgeries, classifying susceptibility to and progression rates of chronic disease such as diabetes, organ rejection, atherogenesis, and venous insufficiency; extreme interventions in trauma cases of particularly high risk and activation-lowering therapies. Also provided is composition derived from a pancreatic homogenate that contains circulating cell activating factors, which can serve as targets for drug screening to identify drug candidates for use in activation lowering therapies. Methods for lowering cell activation by administering protease inhibitors, particularly serine protease inhibitors, are also provided. Kits for performing the methods are also provided.

9003-99-0, Peroxidase IT

RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (methods of diagnosis and triage using cell activation measures)

9003-99-0 CAPLUS ŖΝ

Peroxidase (9CI) (CA INDEX NAME) CM

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

L26 ANSWER 16 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1999:231552 CAPLUS

DOCUMENT NUMBER:

130:249107

TITLE:

System and method for measuring hydrogen peroxide levels in a fluid and method for assessing oxidative

INVENTOR(S): PATENT ASSIGNEE(S): Lacy, Fred; Schmid-Schonbein, Geert W.; Gough, David The Regents of the University of California, USA

SOURCE:

PCT Int. Appl., 39 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:
FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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PATENT NO.
                       KIND
                              DATE
                                        APPLICATION NO.
                                                               DATE
                              ------
                                         _____
                                                               _____
    _____
                       _ _ _ _
                       A1 19990401 WO 1998-US19013 19980914 <--
    WO 9915891
        W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
            DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG,
            KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
            NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
            UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
            FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
            CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
    AU 9894805
                       Α
                              19990412
                                       AU 1998-94805
                                                               19980914 <--
                                                          P 19970925
W 19980914
PRIORITY APPLN. INFO.:
                                         US 1997-60010P
                                         WO 1998-US19013
```

The detection system includes a pair of electrochem. hydrogen peroxide AB sensors, each sensor having working, counter and reference electrodes. A bias voltage is applied to maintain a voltage difference between the working and reference electrodes. A sample aliquot of fluid was treated with either sodium azide or catalase. The sensors are placed in containers containing sufficient amts. of treated fluid to cover the active portions of the electrodes. The output current of each sensor is amplified, and the resulting amplified signals are combined and subtracted to provide a signal which is representative of the level of hydrogen peroxide in the fluid. In a method for assessing oxidative stress, including that related to essential hypertension, the detection system is used to determine a representative level of hydrogen peroxide in blood plasma drawn from a test subject. The level of hydrogen peroxide is directly related to the level of reactive oxygen species in the plasma, and can be used as an accurate predictor of risk for essential hypertension or other conditions related to oxidative stress. Blood plasma samples of normotensive subjects and patients with essential hypertension were analyzed by the system. When hypertensives were compared with family history neg. normotensives, it was found that the hypertensive group had a higher mean arterial pressure by 23% as well as higher levels of plasma hydrogen peroxide by 48% over the normotensive control.

IT 9003-99-0, Myeloperoxidase

RL: ARU (Analytical role, unclassified); ANST (Analytical study) (stabilizer for inhibiting blood catalase and; system and method for measuring hydrogen peroxide levels in fluids and method for assessing oxidative stress)

RN 9003-99-0 CAPLUS

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT:

4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 17 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1999:78214 CAPLUS

DOCUMENT NUMBER:

131:296100

TITLE:

Improved linkage map and thirty new microsatellite

markers for rat Chromosome 10

AUTHOR (S):

Dukhanina, Oksana I.; Sverdlov, Vladimir E.; Hoebee,

Barbara; Rapp, John P.

CORPORATE SOURCE:

Department of Physiology and Molecular Medicine, Medical College of Ohio, Toledo, OH, 43614-5804, USA

SOURCE:

Mammalian Genome (1999), 10(1), 26-29

CODEN: MAMGEC; ISSN: 0938-8990

PUBLISHER: Springer-Verlag New York Inc.

DOCUMENT TYPE: Journal LANGUAGE: English

AB An improved linkage map for rat Chromosome (Chr) 10 with two F2 populations was constructed. Thirty new microsatellite markers were generated from a Chr 10-specific, small-insert genomic library and mapped to rat Chr 10. Among them were the rat homologs for the mouse gene for light and heavy chains of myeloperoxidase and human neurofibromatosis 1. Eight newly generated markers (D10Mco62, D10Mco63, D10Mco64, D10Mco65, D10Mco67, D10Mco68, D10Mco70, and D10Mco74) were mapped to the region of the rat Chr 10 blood pressure QTL. The availability of such markers may be instrumental in the search for genes responsible for the hypertension.

IT 9003-99-0, Myeloperoxidase

RL: BSU (Biological study, unclassified); BIOL (Biological study) (light and heavy chains; improved linkage map and thirty new microsatellite markers for rat chromosome 10: microsatellite sequence similar to intron 7 of light and heavy chains of myeloperoxidase)

RN 9003-99-0 CAPLUS

SOURCE:

in

and

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT: 23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 18 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1998:313962 CAPLUS

DOCUMENT NUMBER: 129:38295

TITLE: New competitive enzyme-linked immunosorbent assay for

determination of metallothionein in tissue and sera

AUTHOR(S): Apostolova, Margarita; Nachev, Choudomir; Koleva,

Milena; Bontchev, Panayot R.; Kehaiov, Ivan

CORPORATE SOURCE: Department of Chemistry and Biochemistry, Medical

Academy, Sofia, 1431, Bulg. Talanta (1998), 46(2), 325-333 CODEN: TLNTA2; ISSN: 0039-9140

PUBLISHER: Elsevier Science B.V.

DOCUMENT TYPE: Journal LANGUAGE: English

AB Very little information is available concerning the relationship between metallothionein (MT) and diseases in humans. Several methods to measure MT levels exist but many of these assays are not sensitive to measure MT in human sera. A new sensitive competitive ELISA system has been developed using MT labeled with horseradish peroxidase as a conjugate and high-titer polyclonal antibodies obtained from rabbit IqG for MT determination

human sera. The cELISA proposed here permits a reliable determination of MT in the range 10-2 000 000 pg ml-1. The method was compared with Cd-hem assay and showed good agreement of results. The recovery of the assay was determined by spiking rat MT into rat and human sera, and comparing it with spiked diluent controls. The overall recoveries of the added MT were 101% for rat sera and 89% for human sera. The variation within-assay and between assay were 3 and 6%, resp. A significant difference (P < 0.001) was found between the MT-level in human sera from patient with essential hypertension $(646\pm223 \text{ ng ml-1}, n = 90)$ and normotensive subjects $(21\pm18 \text{ ng ml-1}, n = 236)$. A correlation between arterial hypertension and MT-level seems possible. A very sensitive new cELISA method was presented for determination of MT in sera and tissues. It enables investigation of possible correlations between sera MT-concentration

certain diseases.

IT 9003-99-0, Peroxidase

RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (horseradish; new competitive ELISA for determination of metallothionein in

tissue and sera)

RN9003-99-0 CAPLUS

Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT: 35 THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 19 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

1998:231271 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 128:253811

TITLE: Expression constructs for the manufacture of

therapeutic proteins as fusion products cleavable by

proteinases manufactured by diseased tissues

Heidtmann, Hans Heinrich; Mueller, Rolf; Sedlacek, INVENTOR(S):

Hans-Harald

Hoechst A.-G., Germany PATENT ASSIGNEE(S):

SOURCE:

Ger., 28 pp.

CODEN: GWXXAW

DOCUMENT TYPE:

Patent German

LANGUAGE: FAMILY ACC. NUM. COUNT:

PATENT INFORMATION: DAMENTO NO

| PA' | TENT NO. | | | | APPLICATION NO. | DATE |
|---------|-----------|--------|--------|-------------|---------------------|-----------------|
| | | | | | | |
| DE | 19701141 | | C1 | 19980409 | DE 1997-19701141 | 19970116 < |
| CN | 1192473 | | A | 19980909 | CN 1998-103791 | 19980114 < |
| EP | 859058 | | A2 | 19980819 | EP 1998-100632 | 19980115 < |
| EP | 859058 | | A3 | 20030528 | | |
| | R: AT, | BE, C | H, DE, | DK, ES, FR, | GB, GR, IT, LI, LU, | NL, SE, MC, PT, |
| | IE, | | | | | , , , , |
| HU | 9800075 | | A2 | 19981028 | HU 1998-75 | 19980115 < |
| RU | 2204414 | | . C2 | 20030520 | RU 1998-100678 | 19980115 |
| CA | 2227159 | | A1 | 19980716 | CA 1998-2227159 | 19980116 < |
| AU | 9852107 | | Α | 19980723 | AU 1998-52107 | 19980116 < |
| AU | 738717 | | B2 | 20010927 | | |
| JP | 10210973 | | Α | 19980811 | JP 1998-6589 | 19980116 < |
| BR | 9800341 | | Α | 19990629 | BR 1998-341 | 19980116 < |
| US | 6080575 | | Α, | 20000627 | US 1998-8308 | 19980116 < |
| US | 6670147 | | B1 | 20031230 | US 1999-256237 | 19990224 |
| US | 200411068 | 32 | A1 | 20040610 | US 2003-638537 | 20030812 |
| PRIORIT | Y APPLN. | INFO.: | | | DE 1997-19701141 | A 19970116 |
| | | | | | US 1998-8308 | A3 19980116 |
| | | | | | US 1999-256237 | A3 19990224 |

- AB A method of manufacturing therapeutically useful proteins as fusion products that can be cleaved and activated in situ by proteinases secreted by diseased or damaged tissue is described. The therapeutic protein is manufactured as a fusion protein with a protein that inhibits its therapeutic action with the two moieties connected by a peptide that is a substrate for a proteinase that is present at increased levels in the diseased tissue. The expression constructs may use tissue-specific promoters to drive expression of the chimeric gene and further increase the specificity of the treatment.
- 9003-99-0D, Peroxidase, fusion products RL: BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (in situ proteolytic activation of; expression constructs for manufacture of therapeutic proteins as fusion products cleavable by proteinases manufactured by diseased tissues)
- RN 9003-99-0 CAPLUS
- Peroxidase (9CI) (CA INDEX NAME) CN

^{***} STRUCTURE DIAGRAM IS NOT AVAILABLE ***

L26 ANSWER 20 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1998:197390 CAPLUS

DOCUMENT NUMBER:

128:253008

TITLE:

Pharmaceutical compositions and methods using alcohols and analogs thereof for regulation of melanin content

and treatment of skin and other diseases

INVENTOR(S):

Brown, David A.; Khorlin, Alexander A.; Lesiak,

Krystyna; Ren, Wu Yun

PATENT ASSIGNEE(S):

Codon Pharmaceuticals, Inc., USA

SOURCE:

PCT Int. Appl., 100 pp.

•

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| | TENT NO. | | | KINI | | DATE | | | APPL: | ICAT | ION 1 | NO. | | D | ATE | |
|---------|--------------------------|-------------------|-----|------------------|------------|--------------------|--------------------|------------|-------|------------|------------|------------|-----|-------------|------------|-------|
| | 9811882 W: AL, KP, | AU, KR, SK, | LC, | A1 BB, LK, | BG, LR, | 1998 BR, LT, | 0326 CA, LV, | CN, MG, | MK, | CZ, MN, | EE, MX, | GE, NO, | NZ, | IL, PL, | IS, RO, | SG, |
| | RW: GH, | | ΙE, | IT, | LU, | MC, | NL, | | | | | | | | | |
| CA | 2266496 | , | , | A1 | | 1998 | | | CA 1 | 997- | 2266 | 496 | | 1 | 9970 | 918 < |
| | 9745842 | | | A | | 1998 | | | AU 1: | | | | | | | 918 < |
| | 740783 | | | B2 | | 2001 | | | AU 1. | , , | 1501. | _ | | _ | ,,, | J10 \ |
| | 5990177 | | | A | | 1999 | | | US 1: | 997_ | 9221 | 4.4 | | 1 | 9970 | 918 < |
| | 957903 | | | A1 | | 1999 | | | EP 1: | 997- | 9443 | 19 | | 1 | | 918 < |
| | 957903 | | | B1 | | 2005 | | | DF I. | J | J44J. | 1.5 | | _ | 2210 | J10 < |
| EF | R: AT, | BE, FI | CH, | | | | | GB, | GR, | IT, | LI, | LU, | NL, | SE, | MC, | PT, |
| US | 6110975 | | | Α | | 2000 | 0829 | | US 1 | 997- | 93314 | 45 | | 1 | 9970 | 918 < |
| AT | 301459 | | | T | | 2005 | 0815 | | AT 1 | 997- | 9443 | 19 | | 1 | 9970 | 918 |
| ES | 2245465 | | | Т3 | | 2006 | 0101 | | ES 1: | 997- | 9443 | 19 | | 1 | 9970 | 918 |
| WO | 9855085 | | | A1 | | 1998 | 1210 | | WO 1 | 998-1 | US534 | 46 | | 1 | 9980 | 318 < |
| | W: AL, | AU, | BA, | BB, | | | | | | | | | HU, | IL, | IS, | JP, |
| | | KR, | | | | | | | | | | | | | | |
| | | SK, | | | | | | | | | | | | | | |
| | | TJ, | | | | , | • | • | • | , | | | | • | • | • |
| | RW: GH, | GM. | KE, | LS, | MW, | SD, | SZ, | UG, | ZW, | AT, | BE, | CH, | DE, | DK, | ES, | FI, |
| | | GB, | | | | | | | | | | | | | | |
| | | GN, | | | | | | | • | • | • | • | | • | • | , |
| AU | 9865659 | • | • | A | | 1998 | | | AU 19 | 998- | 6565 | 9 | | 1 | 9980 | 318 < |
| | 6214888 | | | В1 | | 2001 | | | US 1 | 998- | 8654 | 7 | | 1 | 9980 | 528 < |
| | 6290937 | | | B1 | | 2001 | | | US 1 | 998- | 8591 | 7 | | | | 528 < |
| | 20021419 | 52 | | A1 | | 2002 | | | | | | | | | | |
| | 6623724 | | | B2 | | 2003 | | | | | | | | | | |
| | 20040672 | 0.9 | | A1 | | 2004 | | | US 20 | 003- | 6676 | 3.0 | | 2 | 0030 | 922 |
| | 6955804 | •- | | B2 | | 2005 | | | - | | | | | _ | | |
| | 20061209 | 76 | | | | 2006 | | | US 20 | 005- | 2512 | 1.7 | | 2 | 0051 | 014 |
| | APPLN. | | | ••• | | | | | US 1 | | | | | P 1 | | |
| INTONIT | | 1111 0 . | • | | | | | | US 19 | | | | | P 1 | | |
| | | | | | | | | | US 19 | | | | | | 9970 | |
| | | | | | | | | | US 19 | | | | | | 9970 | |
| | | | | | • | | | | US 1: | | | | | B2 1 | | |
| | • | | | | | | | | WO 19 | | | | | | 9970 | |
| | | | | | | | | | WO 1 | | | | | | 9980 | |
| | | | | | | | | | US 19 | - | | | | w 1 Al 1 | | |
| | | | | | | | | | 00 1. | 790- | UJJI | , | | UT T | J J U U | 220 |

OTHER SOURCE(S): MARPAT 128:253008

Disclosed are methods and compns. for regulating the melanin content of mammalian melanocytes; regulating pigmentation in mammalian skin, hair, wool or fur; treating or preventing various skin and proliferative disorders; increasing the differentiation of mammalian neuronal cells for purposes of treating neurodegenerative diseases or nerve damage; and stimulating cellular nitric oxide (NO) synthesis, cyclic guanosine monophosphate levels (cGMP), and protein kinase G (PKG) activity for purposes of treating diseases mediated by deficiencies in the NO/cGMP/PKG signal transduction pathway; by administration of various compds., including alcs., diols and/or triols and their analogs.

IT 9002-10-2, Tyrosinase

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(alcs. and analogs for regulation of melanin content and treatment of skin and other diseases)

RN 9002-10-2 CAPLUS

AUTHOR (S):

CN Oxygenase, monophenol mono- (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 21 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1998:107525 CAPLUS

DOCUMENT NUMBER: 128:151918

TITLE: Morphological and phytochemical investigations on

Crataegus curvisepala and Crataegus oxyacantha Ghassemi Dehkordi, N.; Ghannadi, A. R.; Mohtaj, F.

CORPORATE SOURCE: Phamacognosy Department, Faculty of Pharmacy and

Pharmaceutical Sciences, Isfahan University of Medical

Sciences, Iran

SOURCE: Daru, Journal of the School of Pharmacy, Tehran

University of Medical Sciences and Health Services (

1996), 6(1&2), 25-36 Persian

CODEN: DJPSFS

PUBLISHER: Tehran University of Medical Sciences, School of

Pharmacy

DOCUMENT TYPE: Journal LANGUAGE: Persian

AB Several species of the genus Crataegus have been used for treatment of hypertension and certain cardiac disorders. In this study, C. curvisepala was examined botanically and phytochem. in comparison to C. oxyacantha. Morphol. and microscopic characteristics of C. curvisepala were examined and some differences were notes from C. oxyacantha. By means of TLC, rutin, hyperoside and chlorogenic acid were identified in these plants.

IT 327-97-9, Chlorogenic acid

RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence)

(morphol. and phytochem. investigations of Crataegus curvisepala and Crataegus oxyancantha)

RN 327-97-9 CAPLUS

CN Cyclohexanecarboxylic acid, 3-[[3-(3,4-dihydroxyphenyl)-1-oxo-2-propenyl]oxy]-1,4,5-trihydroxy-, (1S,3R,4R,5R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

L26 ANSWER 22 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1997:600476 CAPLUS

DOCUMENT NUMBER:

127:253196

TITLE:

Use of (E)-1-(4-(2-alkylaminoethoxy)phenyl)-1-(3-hydroxyphenyl)-2-phenylbut-1-enes for inhibiting

pathological conditions

INVENTOR(S):

Maclean, David Burton; Thompson, David Duane

PATENT ASSIGNEE(S):

Pfizer Inc., USA

SOURCE:

Eur. Pat. Appl., 28 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|------------------------|--------|-----------|------------------------|------|--------------|
| | | | | - | |
| EP 792640 | A2 | 19970903 | EP 1997-301149 | | 19970221 < |
| EP 792640 | A3 | 19980708 | | | |
| R: AT, BE, CH, | DE, DK | , ES, FI, | FR, GB, GR, IE, IT, LI | , LU | , NL, PT, SE |
| US 5985932 | A | 19991116 | US 1997-804346 | | 19970221 < |
| CA 2198571 | A1 | 19970828 | CA 1997-2198571 | | 19970226 < |
| AU 9714956 | A | 19970904 | AU 1997-14956 | | 19970226 < |
| AU 707455 | B2 | 19990708 | | | |
| ZA 9701710 | A | 19980827 | ZA 1997-1710 | | 19970227 < |
| CN 1165651 | A | 19971126 | CN 1997-103416 | | 19970228 < |
| JP 09328421 | A | 19971222 | JP 1997-45616 | | 19970228 < |
| PRIORITY APPLN. INFO.: | | | US 1996-12401P | P | 19960228 |
| | | | US 1996-12402P | P | 19960228 |
| | | | US 1996-12403P | P | 19960228 |
| | | | US 1996-12404P | P | 19960228 |
| | | | US 1996-12410P | P | 19960228 |
| | | | US 1996-12411P | P | 19960228 |
| | | | | | |

OTHER SOURCE(S): MARPAT 127:253196

AB (E)-1-(4-(2-alkylaminoethoxy)phenyl)-1-(3-hydroxyphenyl)-2-phenylbut-1enes are used for the manufacture of a medicament for inhibiting a condition
selected from pathol. conditions related to organ systems which respond to
estrogen agonists, uterine fibrosis, myeloperoxidase activity, autoimmune
diseases, reperfusion damage in ischemic myocardium, and the symptoms of
premenstrual syndrome. An example compound is droloxifene and a number of
pharmaceutical formulations were given.

IT 9003-99-0, Myeloperoxidase

RL: BSU (Biological study, unclassified); BIOL (Biological study) (inhibition of; (alkylaminoethoxyphenyl) (hydroxyphenyl) phenylbutenes for inhibiting pathol. conditions)

RN 9003-99-0 CAPLUS

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

L26 ANSWER 23 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

1997:475087 CAPLUS ACCESSION NUMBER:

127:113366 DOCUMENT NUMBER:

Calcium mineral-based biodegradable microparticles TITLE:

Nuwayser, Elie S. INVENTOR(S): PATENT ASSIGNEE(S): Biotek, Inc., USA U.S., 13 pp. SOURCE: CODEN: USXXAM

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|---------------------|----------|---------------|-------------------------|--------------------|
| | | A | 19970715 | US 1995-538635 | |
| PRIC | RITY APPLN. INFO.: | | | US 1995-538635 | |
| AB | A novel method of p | roduci | ng biodegrada | able microparticles is | disclosed. |
| | Inorg, calcium salt | s are n | nixed with wa | ater to form a slurry. | The slurry is |
| | _ | | | en mixed to form an emu | - |
| | | | | sufficient to form har | |
| | | | | cticles are then retrie | |
| | | | | | |
| | | | _ | ay be added to the slur | |
| | | | | to the hardened micropa | |
| | production The mid | roparti | icles may be | injected into a human | being whereby they |
| | act as controlled-r | elease | drug deliver | ry vehicles. Calcium s | ulfate |
| | hemihydrate (prepar | ed by h | neating calci | um sulfate dihydrate a | t 130° |
| | | | | g, and water 8 mL were | |
| | | | | 00 mL of corn oil at ro | |
| | | | | | |
| | | | | opm for 30 min. The sp | |
| | | | | nin then the microparti | cles were |
| | isolated, washed, a | and drie | ed. | | |
| TITO | 0000 00 0 0 | | | | |

9003-99-0, Peroxidase ΤT

> RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (calcium mineral-based biodegradable microparticles)

9003-99-0 CAPLUS RN

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

L26 ANSWER 24 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1997:464446 CAPLUS

DOCUMENT NUMBER: 127:134146

TITLE:

LTB4-induced accumulation of neutrophils in the lung

plays a role in monocrotaline-induced pulmonary

hypertension

Tabata, Toshiharu AUTHOR(S):

CORPORATE SOURCE: Inst. Dev., Aging Cancer, Tohoku Univ., Sendai,

980-77, Japan

SOURCE: Karei Igaku Kenkyusho Zasshi (1997),

48(3/4), 147-159

CODEN: KIKZEP; ISSN: 1340-3397

PUBLISHER: Tohoku Daigaku Karei Igaku Kenkyusho Kenkyukai

DOCUMENT TYPE: Journal LANGUAGE: Japanese

Monocrotaline (MCT) causes lung inflammation and chronic pulmonary hypertension associated with lung vascular remodeling in rats. We hypothesized that leukotriene B4 (LTB4)-induced accumulation of neutrophils in the lung plays a role in MCT-induced lung disease, and therefore measured LTB4 and myeloperoxidase (MPO) levels in the lung tissue of MCT-treated rats at first. Next, we examined the effect of either specific LTB4 receptor antagonists (ONO4057 or SM15178) or neutropenia (induced by vinblastine sulfate or monoclonal antineutrophils antibody

(RP-3)) on the development of pulmonary hypertension induced by MCT. Lung LTB4 and MPO levels increased at 3 days after MCT injection. In the ONO4057 or Vinblastine treated MCT rats, lung MPO levels were significantly lower than those of MCT-treated rats. At 3 wk after MCT injection, it had caused increases in mean pulmonary arterial pressure, the ratio of right ventricular weight to left ventricle + septum weight (RV/[LV + S]), and the media wall thickness of the muscular arteries of the lung. Treatment with ONO4057 or SM15178, either for 3 wk or during the first week after MCT injection, significantly ameliorated these structural changes. Also, neutropenic rats (induced by either Vinblastine or RP-3) showed significantly lower pulmonary arterial pressure, RV/(LV + S) ratio and the media wall thickness when compared with those of non-treated MCT These results indicate that these LTB4 antagonists inhibited the development of pulmonary hypertension induced by MCT and suggest a role for neutrophils accumulated in the lung tissue in the inflammatory process that contributes to the development of pulmonary hypertension of MCT treated rats. 9003-99-0, Myeloperoxidase

TΤ

RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence)

(LTB4-induced accumulation of neutrophils in lung role in monocrotaline-induced pulmonary hypertension)

RN9003-99-0 CAPLUS

CNPeroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

CAPLUS COPYRIGHT 2007 ACS on STN L26 ANSWER 25 OF 57

ACCESSION NUMBER:

1997:194106 CAPLUS

DOCUMENT NUMBER:

126:262555

TITLE:

AUTHOR(S):

Protective role of synthetic sialylated

oligosaccharide in sepsis-induced acute lung injury Ridings, Philip C.; Holloway, Sharon; Bloomfield, Geoffrey L.; Phillips, M. L.; Fisher, Bernard J.;

Blocher, Charles R.; Sugerman, Harvey J.; Fowler,

Alpha A., III

CORPORATE SOURCE:

Department of Surgery, Medical College of Virginia, Virginia Commonwealth University, Richmond, VA, 23298,

USA

SOURCE:

Journal of Applied Physiology (1997), 82(2),

644-651

CODEN: JAPHEV; ISSN: 8750-7587 American Physiological Society

DOCUMENT TYPE:

PUBLISHER:

LANGUAGE:

Journal English

Proper engagement of leukocyte and endothelial cell selectins with their counterreceptors is an initial step in neutrophil trafficking to sites of inflammation. Certain fucosylated carbohydrate determinants such as sialyl Lewis-x are proposed to act as these counterreceptors. We studied the effects of a synthetic sialyl Lewis-x analog, CY-1503, on the course. of hemodynamic derangements and acute lung injury during exptl. gram-neg. sepsis. Anesthetized ventilated swine were made septic with an infusion of live Pseudomonas aeruginosa. A treatment group received an initial bolus of CY-1503 (60 mg/kg) before sepsis, followed by continuous infusion of CY-1503 (12 mg \cdot kg-1 \cdot h-1). Treatment with CY-1503 did not prevent the development of pulmonary hypertension, systemic hypotension, decline in cardiac output, or severe neutropenia. However, CY-1503 significantly attenuated lung injury, demonstrated by decreased bronchoalveolar lavage protein content and neutrophil influx, lowered lung myeloperoxidase activity, and improved arterial oxygenation. Neutrophils from septic and CY-1503 animals showed significant activation, reflected by upregulated CD18 expression and priming for oxidant burst compared with control animals. This study suggests blockade of selection interactions as a potential therapeutic intervention in sepsis-induced lung injury.

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9003-99-0, Myeloperoxidase
TT
    RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
    BIOL (Biological study); OCCU (Occurrence)
        (protective role of synthetic sialylated oligosaccharide in
        sepsis-induced acute lung injury)
     9003-99-0 CAPLUS
RN
     Peroxidase (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L26 ANSWER 26 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER: 1997:165206 CAPLUS
DOCUMENT NUMBER:
                       126:154428
                       Process for the identification of proteolytic
TITLE:
                      activities and/or inhibitors thereof
                       Fassina, Giorgio; Corti, Angelo
INVENTOR(S):
                      Tecnogen S.C.P.A., Italy
PATENT ASSIGNEE(S):
SOURCE:
                       Eur. Pat. Appl., 20 pp.
                       CODEN: EPXXDW
DOCUMENT TYPE:
                       Patent
                       English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
                       2
PATENT INFORMATION:
                   KIND DATE
                                       APPLICATION NO.
    PATENT NO.
                                                              DATE
     _____
                       _ _ _ _
                              -----
                                         ______
                                                               _____
    EP 751225
                      A1 19970102
                                       EP 1996-114931
                                                               19911014 <--
    EP 751225
                       B1 20010328
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE
                                                              19911014 <--
    EP 481930 A2 19920422
                                       EP 1991-830428
    EP 481930
                       A3
                             19930630
    EP 481930
                       B1
                             19970618
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE
    AT 154609 T 19970715 AT 1991-830428 19911014 <--
    AT 200107
                        \mathbf{T}
                              20010415
                                         AT 1996-114931
                                                               19911014 <--
                                         IT 1990-48365
PRIORITY APPLN. INFO.:
                                                          A 19901015
                                         IT 1991-RM261
                                                           A 19910415
                                                           A3 19911014
                                         EP 1991-830428
                                         IT 1991-R0261
                                                               19910415
ΔR
    This invention relates to a process for the identification of proteolytic
    activities or of activities that inhibit proteolytic activities,
    particularly of endothelin and/or of TNF, especially in biol. fluids,
fermentation
    broths, conditioned culture soils, cell exts., and plant exts. As an
    example, the process can use a fragment of proendothelin as substrate as
    well as a ligand comprising amino acid sequences that are hydropathically
    complementary to the fragment of proendothelin.
TΤ
    9003-99-0D, Peroxidase, streptavidin conjugates
    RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)
        (determination of proendothelin- and TNF-specific proteolytic activities and
       their inhibitors)
    9003-99-0 CAPLUS
RN
CN
    Peroxidase (9CI) (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L26 ANSWER 27 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER:
                       1997:121111 CAPLUS
DOCUMENT NUMBER:
                       126:135587
TITLE:
                       Extraction of red pigments from apples as
```

antihypertensives, allergy inhibitors,

Shimoda, Shunji

INVENTOR (S):

antioxidants, anticaries agents and deodorants

Tanabe, Masayuki; Kanda, Tomomasa; Yanagida, Akiro;

PATENT ASSIGNEE(S):

Nikka Whisky, Japan

SOURCE:

Jpn. Kokai Tokkyo Koho, 10 pp.

CODEN: JKXXAF

DOCUMENT TYPE:

Patent

LANGUAGE:

FAMILY ACC. NUM. COUNT:

Japanese

PATENT INFORMATION:

| · ; | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|-------|---------------------|---------|--------------|-------------------------|-----------------|
| | | | | | |
| | JP 08319433 | Α | 19961203 | JP 1995-130327 | 19950529 < |
| PRIOR | ITY APPLN. INFO.: | | | JP 1995-130327 | 19950529 |
| | | | | phenols] are extracted | from apples for |
| | | | | bitors, antioxidants, | |
| | anticaries agents a | nd deod | orants (hali | tosis inhibitors). In | vitro expts. |
| : | indicated that the | red pig | ments inhibi | ted the histamine relea | se from |
| (| cultured RBL-2H3 ce | lls, in | dicating ant | iallergy activity. | |

IT 327-97-9P, Chlorogenic acid

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(extraction of red pigments from apples as antihypertensives, allergy inhibitors, antioxidants, anticaries agents and deodorants)

RN 327-97-9 CAPLUS

CN Cyclohexanecarboxylic acid, 3-[[3-(3,4-dihydroxyphenyl)-1-oxo-2propenyl]oxy]-1,4,5-trihydroxy-, (1S,3R,4R,5R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

L26 ANSWER 28 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1996:565344 CAPLUS

DOCUMENT NUMBER:

125:244617

TITLE:

Allopurinol reduces bacterial translocation, intestinal mucosal lipid peroxidation, and

neutrophil-derived myeloperoxidase activity in chronic

portal hypertensive and common bile duct-ligated

growing rats

AUTHOR (S):

Schimpl, Gunther; Pesendorfer, Patricia; Steinwender,

Gerhard; Feierl, Gerhard; Ratschek, Manfred;

Hollwarth, Michael E.

CORPORATE SOURCE:

Medical School, University Graz, A-8036, Austria

Pediatric Research (1996), 40(3), 422-428

CODEN: PEREBL; ISSN: 0031-3998

PUBLISHER:

SOURCE:

Williams & Wilkins

DOCUMENT TYPE:

Journal English

LANGUAGE:

Bacterial translocation (BT) from the gastrointestinal tract has been thought to play a role in the pathogenesis of septic complications in

patients with chronic portal hypertension (PH) and obstructive jaundice. The purpose of this study was to investigate the incidence of BT and to assess the role of intestinal mucosal malondialdehyde (MDA) levels as an indicator of lipid peroxidn. and polymorphonuclear neutrophil-derived myeloperoxidase (MPO) in chronic portal hypertensive and common bile duct-ligated rats. Twenty male rats were subjected to sham laparotomy (SL), 20 rats to calibrated portal vein constriction (PH), 20 rats to common bile duct ligation (CBDL), and 10 rats served as a nonoperated control group (NOP). After 4 wk, 10 animals of each operated group received 50 mg/kg allopurinol i.p., at 24 h, and again 2 h prior to estimation of BT, intestinal mucosal MDA, and MPO activities. In the NOP and SL groups, BT to the mesenteric lymph nodes (MLN) and spleen was present. In PH and in CBDL rats, BT to liver, portal vein, peritoneum, and caval vein occurred. Allopurinol treatment attenuated the frequence of BT in PH and decreased BT in CBDL rats significantly (p < 0.05). Ileal mucosal MDA levels (nanomoles/g) in untreated rats increased from 45,1 ± 7.9 in SL to 98.2 \pm 9.1 in PH and to 102.2 \pm 11 in CBDL rats (p < 0.01). In the allopurinol groups the increase of MDA to 49.1 \pm 1.3 in PH, and 66.2 ± 2.2 in CBDL was significantly lower (p < 0.01). MPO activity (units/g) in the ileal mucosa increased in untreated rats from 319 \pm 129 after SL to 866 \pm 104 after PH and to 1016 \pm 104 after CBDL (p < 0.01). Allopurinol significantly attenuated MPO activity to 369 ± 44 in PH, and to 372 \pm 60 in CBDL animals (p < 0.01). In PH and CBDL rats significant BT, intestinal mucosal lipid peroxidn., and polymorphonuclear neutrophil-derived MPO activity occurred. Allopurinol reduced BT and improved intestinal mucosal MDA and MPO activities, suggesting that there might be an association between BT and intestinal mucosal lipid peroxidn. 9003-99-0, Myeloperoxidase

TT RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

> (neutrophil; allopurinol reduces bacterial translocation, intestinal mucosal lipid peroxidn., and neutrophil-derived myeloperoxidase activity in chronic portal hypertension and common bile duct ligation)

9003-99-0 CAPLUS RN

Peroxidase (9CI) (CA INDEX NAME) CN

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

L26 ANSWER 29 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1996:493848 CAPLUS

DOCUMENT NUMBER:

125:184971

TITLE:

Allopurinol and glutamine attenuate bacterial translocation in chronic portal hypertensive and

common bile duct-ligated growing rats

AUTHOR (S):

Schimpl, G.; Pesendorfer, P.; Steinwender, G.; Feierl,

G.; Ratschek, M.; Hollwarth, M. E.

CORPORATE SOURCE: SOURCE:

Medical School, University Graz, Graz, A-8036, Austria

Gut (1996), 39(1), 48-53 CODEN: GUTTAK; ISSN: 0017-5749

PUBLISHER:

BMJ Publishing Group

DOCUMENT TYPE: LANGUAGE:

Journal English

Spontaneous bacterial infections and septicemia result in morbidity and mortality in patients with portal hypertension and obstructive jaundice. The aim of this study in rats was to investigate the incidence of bacterial translocation in portal hypertension and obstructive jaundice, and to evaluate the effects of allopurinol and glutamine. Rats were subjected to sham laparotomy (SL), portal hypertension (PH) by calibrated stenosis of the portal vein, and common bile duct ligation (CBDL). Animals of each group were either treated with allopurinol (50 mg/kg twice a week), glutamine (1 g/kg/d), and allopurinol and glutamine. After four weeks, significant bacterial translocation in the untreated PH and CBDL rats occurred. Intestinal

mucosal malondialdehyde concns. (MDA), as an indicator for lipid peroxidn., and myeloperoxidase activity (MPO) released from activated neutrophils were also significantly increased (p<0.01). Allopurinol and glutamine in PH and CBDL rats improved bacterial translocation, and decreased MDA and MPO values (p<0 $\!\Sigma$ 01). In conclusion, in PH and CBDL rats significant bacterial translocation, ileal mucosal lipid peroxidn., and neutrophil derived MPO activity occurred. Allopurinol and glutamine significantly reduced bacterial translocation, as well as ileal mucosal MDA and MPO activities.

IT 9003-99-0, Myeloperoxidase

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)

(neutrophil-derived myeloperoxidase; allopurinol and glutamine attenuation of bacterial translocation in chronic portal hypertensive and common bile duct-ligated growing rats)

RN 9003-99-0 CAPLUS

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

L26 ANSWER 30 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1996:336393 CAPLUS

DOCUMENT NUMBER:

125:19009

TITLE:

Solid delivery systems for controlled release of

molecules incorporated therein

INVENTOR(S):

Roser, Bruce Joseph; Colaco, Camilo; Jerrow, Mohamed Abdel Zahra; Blair, Julian Alexander; Kampinga, Jaap;

Wardell, James Lewis; Duffy, John Alistair Quadrant Holdings Cambridge Limited, UK

PATENT ASSIGNEE(S):

PCT Int. Appl., 99 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| | | | | | | | | APPLICATION NO. | | | | | | | | | |
|---------------|------|------|-----|-----|-----------|-------------|-------|-----------------|-----|-------|-------|------|-----|-----|-----|----------|--------|
| | | | | | | A1 19960215 | | | | | | | | | | 9950 | 804 < |
| | W: | AM, | ΑT, | AU, | BB, | BG, | BR, | BY, | CA, | CH, | CN, | CZ, | DE, | DK, | EE, | ES, | FI, |
| | | GB, | GE, | HU, | IS, | JP, | KE, | KG, | ΚP, | KR, | KZ, | LK, | LR, | LT, | LU, | LV, | MD, |
| | | MG, | MN, | MW, | MX, | NO, | NZ, | PL, | PT, | RO, | RU, | SD, | SE, | SG, | SI, | SK, | ТJ, |
| | | TM, | TT | | | | | | | | | | | | | | |
| | RW: | KΕ, | MW, | SD, | SZ, | ŪĠ, | ΑT, | BE, | CH, | DE, | DK, | ES, | FR, | GB, | GR, | ΙE, | IT, |
| | | LU, | MC, | NL, | PT, | SE, | BF, | ВJ, | CF, | CG, | CI, | CM, | GΑ, | GN, | ML, | MR, | NE, |
| | | SN, | TD, | TG | | | | | | | | | | | | | |
| US | 6290 | 991 | | | | | 2001 | 0918 | • | US 1 | 994 - | 3490 | 29 | | 1: | 9941 | 202 < |
| CA | 2197 | 982 | | | A1 | | 1996 | 0215 | | CA 1: | 995-: | 2197 | 982 | | 1: | 9950 | 804 < |
| | 9531 | | | | | | 1996 | | | AU 1: | 995- | 3185 | 1 | | 1 | 9950 | 804 < |
| | 6885 | | | | | | | | | | | | | | | | |
| ĒΡ | 7737 | 81 | | | A1 | | 1997 | 0521 | | EP 1: | 995- | 9278 | 56 | | 1: | 9950 | 804 < |
| ĒΡ | 7737 | 81 | | | B1 | | 2003 | 1022 | | | | | | | | | |
| | | | | | | | | | | | | | | | | | PT, SE |
| JP | 1050 | 3769 | | | ${f T}$ | | 1998 | 0407 | 1 | JP 1: | 995- | 5063 | 45 | | 1: | 9950 | 804 < |
| HU | 7777 | 7 | | | A2 | | 1998 | 0828 | , : | HU 1: | 998- | 594 | | | 1: | 9950 | 804 < |
| | 1204 | | | | | | | | | | | | | | | 9950 | 804 < |
| ΕP | 1138 | 319 | | | A2 | | 2001 | 1004 | | EP 2 | 001- | 1166 | 37 | | 1: | 9950 | 804 < |
| \mathbf{EP} | 1138 | | | | | | 2003 | | | | | | | | | | |
| | R: | ΑT, | ΒE, | CH, | DE, | DK, | ES, | FR, | GB, | GR, | IT, | LI, | LU, | NL, | SE, | MC, | PT, |
| | | | | LT, | | | | | | | | | | | | | |
| EΡ | 1138 | 337 | | | A2 | : | 2001: | 1004 | • | EP 20 | 001- | 1166 | 38 | | 19 | 9950 | 804 < |
| ΕP | 1138 | 337 | | | A3 | : | 2003 | 0326 | | | | | | - | | | |
| | R: | AΤ, | BE, | CH, | DE, | DK, | ES, | FR, | GB, | GR, | IT, | LI, | LU, | NL, | SE, | MC, | PT, |
| | | ΙE, | SI, | LT, | LV | | | | | | | | | | | | |

| RU | 2177785 | C2 | 20020110 | RU | 1997-103529 | | 19950804 | |
|---------------|--------------------|------------|--------------|--------|---------------|----------|------------|-----|
| EE | 3593 | B1 | 20020215 | EE | 1997-62 | | 19950804 | |
| \mathtt{PL} | 184068 | B1 | 20020830 | PL | 1995-318898 | | 19950804 | |
| SK | 283026 | B6 | 20030204 | SK | 1997-277 | • | 19950804 | |
| AT | 252373 | T | 20031115 | AT | 1995-927856 | | 19950804 | |
| PT | 773781 | ${f T}$ | 20040331 | PT | 1995-927856 | | 19950804 | |
| ES | 2208687 | T3 | 20040616 | ES | 1995-927856 | | 19950804 | |
| EP | 1516615 | A2 | 20050323 | | 2004-29125 | | 19950804 | |
| | R: AT, BE, CH, | DE, | DK, ES, FR, | GB, GI | R, IT, LI, LU | , NL, SE | E, MC, PT, | IE |
| CZ | 297431 | В6 | 20061213 | CZ | 1997-476 | | 19950804 | |
| FI | 9700867 | Α | 19970408 | FI | 1997-867 | | 19970228 | |
| | 9701688 | Α | 19970411 | | 1997-1688 | | 19970411 | |
| | 9871864 | Α | 19980820 | AU | 1998-71864 | | 19980612 | < |
| | 707605 | B2 | 19990715 | | | | | |
| | 6331310 | B1 | 20011218 | | 2000-628380 | | 20000801 | |
| | 2001038858 | A1 | 20011108 | US | 2001-755737 | | 20010105 | < |
| | 6586006 | B2 | 20030701 | | | | | |
| | 2002012687 | A1 | 20020131 | US | 2001-945180 | | 20010831 | |
| | 6565871 | B2 | 20030520 | | | | | |
| | 2003054040 | A1 | 20030320 | US | 2002-280468 | | 20021025 | |
| | 6811792 | B2 | 20041102 | | | | | |
| | 2003147961 | A1 | 20030807 | US | 2003-376136 | | 20030227 | |
| | 6893657 | B2 | 20050517 | | | | | |
| | 2004052825 | A1 | 20040318 | US | 2003-652212 | | 20030829 | |
| | 7056495 | B2 | 20060606 | | | | | |
| | 2004219206 | A1 | 20041104 | | 2004-857100 | | 20040528 | |
| | 2005276845 | A1 | 20051215 | | 2005-134573 | | 20050520 | |
| | 2005276846 | A 1 | 20051215 | | 2005-134700 | | 20050520 | |
| | 2005276759 | A1 | 20051215 | | 2005-134701 | | 20050520 | |
| | 2006056898 | Α | 20060302 | | 2005-284596 | | 20050929 | |
| PRIORIT | Y APPLN. INFO.: | | | | 1994-15810 | | 19940804 | |
| | | | | | 1994-349029 | | 19941202 | |
| | | | | | 1995-927856 | | 19950804 | |
| | | | | | 1996-506345 | | 19950804 | |
| | | | | | 1995-GB1861 | W | 19950804 | |
| | | | | | 1997-500877 | | 19970818 | |
| | | | | | 2000-628380 | | 20000801 | |
| | | | | | 2001-116638 | | 20010713 | |
| | | | | | 2001-945180 | | 20010831 | |
| | | | | | 2003-376136 | | 20030227 | |
| | | | | | 2003-652212 | | 20030829 | |
| AB So. | lid dosage deliver | TV SV | stems suitab | le for | delivery of | hioacti | ve materi | ale |

AB Solid dosage delivery systems suitable for delivery of bioactive materials s.c., intradermal, i.m., and i.v. are disclosed. The delivery systems comprise a vitreous vehicle, e.g. polyol, loaded with the guest substance and capable of releasing the guest substance in situ at various controlled rates. Microparticles were prepared by spray drying a solution of 0.39 M trehalose, 0.14 M calcium lactate and 0.5% MB9. This particles were coated by addition of a saturated solution of zinc palmitate in toluene and cooling

at 60-30°. The particles were then filtered under vacuum to remove excess zinc palmitate, washed with acetone, and air-dried. The resulting powder remained unwetted in water for \geq 3 days and released MB9 slowly into the water.

IT 9003-99-0, Peroxidase

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (controlled-release solid delivery systems comprising polyols)

RN 9003-99-0 CAPLUS

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

L26 ANSWER 31 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 1996:166955 CAPLUS

DOCUMENT NUMBER:

124:257280

Characteristics of renal tubular atrophy in TITLE:

experimental renovascular hypertension: a

model of kidney hibernation

Groene, H.-J.; Warnecke, E.; Olbricht, C. J. AUTHOR(S):

Medizinisches Zentrum fur Pathologie, Universitat CORPORATE SOURCE:

Marburg, Marburg/Lahn, D-35043, Germany

Nephron (1996), 72(2), 243-52 SOURCE:

CODEN: NPRNAY; ISSN: 0028-2766

Karger PUBLISHER: DOCUMENT TYPE: Journal English LANGUAGE:

The inability to sep. irreversible lesions of tubular epithelia from reversible tubular atrophy constitutes a major problem in histopathol. and in decisions for revascularization of shrunken kidneys with renal artery stenosis. To characterize reversible tubular atrophy ("kidney hibernation") the authors studied the physiol. and biochem. parameters and morphol. including histochem. in rat kidneys made atrophic by renal artery stenosis and treatment with the angiotensin-converting enzyme inhibitor, enalapril. Renal artery stenosis was induced by a 0.2-mm clip around the left renal artery. Following 7 wk of clipping and 2 concomitant weeks of enalapril treatment, the kidney length decreased from 17.8 to 13.7 mm. Renal blood flow and glomerular filtration rate decreased to 39% and to approx. 3% of control values, resp. The activities of the intracellular proteolytic enzymes cathepsin B and L and of Na-K-ATPase in microdissected proximal tubular segments decreased to values below 50 and 10%, resp. All changes were significant. Histochem. staining for ATPase activity in the distal tubule segments remained unchanged. Tubular cells were atrophic but not necrotic. Histochem. staining of alkaline phosphatase in the tubular brush border and of acid phosphatase and peroxidase in lysosomes was greatly reduced. All observed changes were reversible within 2-3 wk following removal of the clip and withdrawal of enalapril either with or without contralateral nephrectomy. Thus, a form of kidney hibernation with readily reversible tubular atrophy has been described. Based on this description it may be possible in consecutive expts. to differentiate between reversible and irreversible tubular atrophy.

9003-99-0, Peroxidase IT

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)

(enzymic characteristics of renal tubular atrophy (kidney hibernation) in renovascular hypertension)

9003-99-0 CAPLUS RN

CN Peroxidase (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

L26 ANSWER 32 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1996:90279 CAPLUS

DOCUMENT NUMBER: TITLE:

124:142540

Properties of circulating leukocytes in spontaneously

hypertensive rats

AUTHOR (S):

Shen, K.; Sung, K.-L. P.; Whittemore, D. E.; DeLano,

F. A.; Zweifach, B. W.; Schmid-Schoenbein, G. W.

CORPORATE SOURCE:

Dept. Bioengineering, Univ. California, San Diego, CA,

92093-0412, USA

SOURCE:

PUBLISHER:

Biochemistry and Cell Biology (1995), 73(7 &

8), 491-500

CODEN: BCBIEQ; ISSN: 0829-8211 National Research Council of Canada

DOCUMENT TYPE:

LANGUAGE:

Journal English

The factors responsible for predisposition to progressive organ injury and vascular complications in arterial hypertension are uncertain. Recent evidence shows that leukocytes participate in cardiovascular conditions for which hypertension is a risk factor. Therefore,

there is a need to define the properties of circulating leukocytes in hypertensives. There are about twice as many circulating leukocytes in spontaneous hypertensive rats (SHRs) compared with their normotensive controls, the Wistar-Kyoto rats (WKYs). The SHR neutrophils are viscoelastic and similar to neutrophils in WKYs but exhibit lower deformability in short-term elastic deformation. Mature SHRs have elevated levels of spontaneous pseudopod formation. Mild stimulation with N-formyl-Met-Leu-Phe or platelet-activating factor (10-8 M) results in a significantly enhanced level of neutrophil pseudopod formation in SHRs but not in WKYs. SHRs exhibit higher levels of spontaneous superoxide formation. Alkaline phosphatase content of individual circulating neutrophils in SHRs is on average lower while plasma levels of alkaline phosphatase in the same samples are elevated in the SHRs. Spontaneous degranulation of SHR neutrophils is also detectable with myeloperoxidase measurements. Such activity of circulating leukocytes poses a significant risk for vascular cytotoxicity in the hypertensive rats.

TT 9003-99-0, Myeloperoxidase

> RL: ADV (Adverse effect, including toxicity); BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence)

(myelo-; properties of circulating leukocytes in spontaneously hypertensive rats)

9003-99-0 CAPLUS RN

Peroxidase (9CI) CN (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

L26 ANSWER 33 OF 57 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1995:731707 CAPLUS

DOCUMENT NUMBER:

123:123135

TITLE:

Extraction of fruit polyphenols and their uses as antioxidant, hypotensive, antimutagenic agent, antiallergic agent and anticariogenic agent.

INVENTOR(S):

Tanabe, Masayuki; Kanda, Tomomasa; Yanagida, Akio Nikka Whisky Distilling Co., Ltd., Japan

PATENT ASSIGNEE(S):

Eur. Pat. Appl., 34 pp.

SOURCE:

CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION: DATENT NO

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|----------------------------|----------------------|----------------------|--|-----|----------------------------------|
| EP 657169 R: AT, | A1 BE, DE, FR, GE | 19950614 B, IT | EP 1994-401669 | - | 19940720 < |
| CA 2128293 CA 2128293 | A1 C | 19950607 | CA 1994-2128293 | | 19940718 < |
| AU 9468996 AU 683892 | A B2 | 19941013 19971127 | AU 1994-68996 | | 19940809 < |
| CN 1121924 CN 1051089 | A B | 19960508 | CN 1994-115048 | | 19940818 < |
| JP 07285876 JP 3521155 | A B2 | 19951031 20040419 | JP 1994-300578 | | 19941205 < |
| JP 200204719 US 5932623 | | 20020212 | JP 2001-190347 US 1995-555729 | | 19941205 19951109 < |
| JP 08259453 US 5994413 | A A | 19961008 19991130 | JP 1996-86859 US 1997-784546 | | 19960409 < |
| JP 200517937 | 3 A | 20050707 | JP 1993-305632 | 73. | 20050204 |
| FRIORITI APPEN. I | NFO.: | | JP 1994-24435 | Α | 19940222 |
| | | | US 1994-278080 JP 1994-300578 JP 2001-190347 | АЗ | 19940720 19941205 19941205 |

The present invention provides a fruit polyphenol obtained by subjecting unripe fruits of Rosaceae to pressing and/or extraction and then purifying the resulting juice or extract and its uses as antioxidant, hypotensive, antimutagenic agent, antiallergic agent and anticariogenic agent. The fruit polyphenol has various physiol. activities, e.g., antioxidant, an ACE-inhibiting, hyaluronidase-inhibiting and GTase-inhibiting activities. Thus, polyphenols were obtained by crushing unripe apples, while adding an appropriate amount of SO2 and pressing using an oil press. Further, the addition of an enzyme followed by centrifugation or filtration and column chromatog. gave polyphenol powder products. The antimutagenic activity of the polyphenol was demonstrated by using Salmonella typhimurium.

IT 327-97-9, Chlorogenic acid

327-97-9, Chlorogenic acid
RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BSU (Biological study, unclassified); THU (Therapeutic use);
BIOL (Biological study); OCCU (Occurrence); USES (Uses)

(fruit polyphenols as pharmaceuticals)

RN 327-97-9 CAPLUS

CN Cyclohexanecarboxylic acid, 3-[[3-(3,4-dihydroxyphenyl)-1-oxo-2-propenyl]oxy]-1,4,5-trihydroxy-, (1S,3R,4R,5R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

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         OCT 30
                CHEMLIST enhanced with new search and display field
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         NOV 03
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NEWS
     6
        NOV 10
                CA/CAplus F-Term thesaurus enhanced
NEWS
         NOV 10
     7
                STN Express with Discover! free maintenance release Version
                 8.01c now available
NEWS
         NOV 20
     8
                CA/CAplus to MARPAT accession number crossover limit increased
                 to 50,000
NEWS 9
        DEC 01
                CAS REGISTRY updated with new ambiguity codes
NEWS 10
         DEC 11
                CAS REGISTRY chemical nomenclature enhanced
         DEC 14
NEWS 11
                WPIDS/WPINDEX/WPIX manual codes updated
NEWS 12
        DEC 14
                GBFULL and FRFULL enhanced with IPC 8 features and
                 functionality
NEWS 13
         DEC 18
                CA/CAplus pre-1967 chemical substance index entries enhanced
                 with preparation role
NEWS 14
         DEC 18
                CA/CAplus patent kind codes updated
NEWS 15
        DEC 18
                MARPAT to CA/Caplus accession number crossover limit increased
                 to 50,000
NEWS 16 DEC 18
                MEDLINE updated in preparation for 2007 reload
NEWS 17
        DEC 27
                CA/CAplus enhanced with more pre-1907 records
NEWS 18
        JAN 08
                CHEMLIST enhanced with New Zealand Inventory of Chemicals
NEWS 19
                CA/CAplus Company Name Thesaurus enhanced and reloaded
         JAN 16
NEWS 20
        JAN 16
                IPC version 2007.01 thesaurus available on STN
NEWS 21 JAN 16
                WPIDS/WPINDEX/WPIX enhanced with IPC 8 reclassification data
NEWS 22
        JAN 22
                CA/CAplus updated with revised CAS roles
NEWS 23
         JAN 22
                CA/CAplus enhanced with patent applications from India
NEWS 24
         JAN 29
                PHAR reloaded with new search and display fields
NEWS 25
        JAN 29
                CAS Registry Number crossover limit increased to 300,000 in
                 multiple databases
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120 L6 NOT FERULIC ACID

=> s 17 not caffeic acid

8629 CAFFEIC

4310472 ACID

1567929 ACIDS

4811583 ACID

(ACID OR ACIDS)

8042 CAFFEIC ACID

(CAFFEIC (W) ACID)

L8 98 L7 NOT CAFFEIC ACID

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PROCESSING COMPLETED FOR L8

98 FOCUS L8 1-

=> d ibib abs 1-98 hitstr

ANSWER 1 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

2006:328632 CAPLUS ACCESSION NUMBER:

TITLE: Study on NO-donating antihypertensive agents

I. synthesis and antihypertensive activity

of C-3 nitrate or furoxan substituted benzopyrans

AUTHOR (S): Xu, Xin; Zhang, Yihua; Peng, Sixun; Ji, Hui; Li,

Yongqi

CORPORATE SOURCE: Center of Drug Discovery, China Pharmaceutical

University, Nanjing, 210009, Peop. Rep. China

SOURCE: Zhongguo Yaoke Daxue Xuebao (2005), 36(6), 488-495

CODEN: ZHYXE9; ISSN: 1000-5048

Zhongguo Yaoke Daxue PUBLISHER:

DOCUMENT TYPE: Journal Chinese LANGUAGE:

The synthesis and antihypertensive activity of NO-donating benzopyran compds. were studied to search for novel antihypertensive agents with better efficacy and fewer side-effects. Sixteen novel compds. were synthesized by coupling of organic nitrate and furoxan with trans-4-(acetyloxy)-3,4-dihydro-3-hydroxy-2,2dimethyl-2H-1-benzopyran-6-carbonitrile and succinic acid. The inhibition of the target compds. on KCl-induced contraction of aortic strips and the effects on systolic aortic pressure (SAP) and diastolic aortic pressure (DAP) of the spontaneously hypertensive rats (SHR) were

measured. The amount of NO released in vitro of the compds. was also

determined

by Griess method. The preliminary pharmacol. testings showed that most of the target compds. inhibited the KCl-induced contraction to some extent. Among them, butanedioic acid 4-acetoxy-6-cyano-2,2-dimethyl-3,4-dihydro-2Hbenzopyran-3-yl 4-(3-phenylsulfonyl-5-oxido-1,2,5-oxadiazol-4-yloxy)butyl ester decreased SAP (15.2%) and PAP (12.5%) of the SHR with a longer duration than did control pinacidil. The amount of NO released by this compound was 0.9 µg/mL. The relationship between NO release and the antihypertensive effect of the target compds. remains to be investigated.

ΙT INDEXING IN PROGRESS

IT1135-24-6, 4-Hydroxy-3-methoxycinnamic acid

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of nitric oxide-releasing antihypertensive agents bearing benzopyran and nitrate or furoxan substituents)

RN 1135-24-6 CAPLUS

CN 2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)- (9CI) (CA INDEX NAME)

L9 ANSWER 2 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2001:567759 CAPLUS

DOCUMENT NUMBER: 135:298179

TITLE: Novel hypotensive agents from Verbesina caracasana. 8.

Synthesis and pharmacology of (3,4-dimethoxycinnamoyl)-

N1-agmatine and synthetic analogues

AUTHOR(S): Carmignani, Marco; Volpe, Anna Rita; Botta, Bruno;

Espinal, Romulo; De Bonnevaux, Stella C.; De Luca, Carlo; Botta, Maurizio; Corelli, Federico; Tafi, Andrea; Sacco, Rosario; Delle Monache, Giuliano

CORPORATE SOURCE: Dipartimento di Biologia di Base e Applicata Sezione

di Farmacologia, Universita di L'Aquila, Coppito (AQ),

67010, Italy

SOURCE: Journal of Medicinal Chemistry (2001), 44(18),

2950-2958

CODEN: JMCMAR; ISSN: 0022-2623

American Chemical Society

DOCUMENT TYPE: Journal LANGUAGE: English

PUBLISHER:

LANGUAGE: English

OTHER SOURCE(S): CASREACT 135:298179

The more polar metabolites from the Venezuelan plant Verbesina caracasana, i.e., N3-prenylagmatine, (3,4-dimethoxycinnamoyl)-N1-agmatine, agmatine, and galegine (prenylguanidine), previously reported (Delle Monache, G.; et al. BioMed. Chemical Lett. 1999, 9, 3249-3254), have been synthesized following a biosynthetic strategy. The pharmacol. profiles of various synthetic analogs of (3,4-dimethoxycinnamoyl)-N1-agmatine (G5) were also analyzed, to shed some light on the structure-activity relationship of these compds. Derivs. with the (E)-configuration and/or with a p-methoxybenzoyl moiety were found to be responsible for higher hypotensive effects, which were associated with a slight and, in some cases, not dose-related increase of cardiac inotropism, with variable and not significant chronotropic responses, and, only at higher doses, with effects of respiratory depression. Either an increase (to six) or a decrease (to two) of the number of methylene groups in the alkyl chain of (E)-G5 did not change blood pressure responses, while slightly increasing the pos. inotropic ones. At pharmacol. doses, all the studied compds. showed hypotensive and slight pos. inotropic effects without relevant chronotropic and respiratory actions.

IT 146072-40-4P 365568-01-0P 365568-02-1P 365568-03-2P 365568-04-3P 365568-05-4P

365568-06-5P 365568-07-6P 365568-08-7P

RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); PROC (Process); RACT (Reactant or reagent)

(design of antihypertensive drugs from Verbesina caracasana)

RN 146072-40-4 CAPLUS

CN

2-Propenamide, N-[4-[(aminoiminomethyl)amino]butyl]-3-(3,4-dimethoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

MeO
$$E$$
 NH
 NH_2
 NH_2
 NH_2

RN 365568-01-0 CAPLUS

CN Carbamic acid, [[4-[[(2E)-3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]butyl]carbonimidoyl]bis-, bis(1,1-dimethylethyl) ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 365568-02-1 CAPLUS

CN Carbamic acid, [[2-[[(2E)-3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]ethyl]carbonimidoyl]bis-, bis(1,1-dimethylethyl) ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 365568-03-2 CAPLUS

CN Carbamic acid, [[3-[[(2E)-3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]propyl]carbonimidoyl]bis-, bis(1,1-dimethylethyl) ester (9CI) (CA INDEX NAME)

RN 365568-04-3 CAPLUS

CN Carbamic acid, [[5-[[(2E)-3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]pentyl]carbonimidoyl]bis-, bis(1,1-dimethylethyl) ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 365568-05-4 CAPLUS

CN Carbamic acid, [[6-[[(2E)-3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]hexyl]carbonimidoyl]bis-, bis(1,1-dimethylethyl) ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 365568-06-5 CAPLUS

CN 2-Propenamide, N-[2-[(aminoiminomethyl)amino]ethyl]-3-(3,4-dimethoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & \\ & & & \\$$

RN 365568-07-6 CAPLUS

CN 2-Propenamide, N-[3-[(aminoiminomethyl)amino]propyl]-3-(3,4-dimethoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

MeO
$$E$$
 NH NH_2 NH_2

RN 365568-08-7 CAPLUS

CN 2-Propenamide, N-[5-[(aminoiminomethyl)amino]pentyl]-3-(3,4dimethoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

MeO
$$\stackrel{E}{\underset{H}{\overset{O}{\longrightarrow}}}$$
 $\stackrel{NH}{\underset{H}{\overset{NH}{\longrightarrow}}}$ $\stackrel{NH}{\underset{H}{\overset{NH}{\longrightarrow}}}$

IT 128009-16-5, 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4 [[imino[(3-methyl-2-butenyl)amino]methyl]amino]butyl] RL: ADV (Adverse effect, including toxicity); BPR (Biological process);
 BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
 (Biological study); PROC (Process); USES (Uses)
 (design of antihypertensive drugs from Verbesina caracasana)
RN 128009-16-5 CAPLUS
CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[[imino[(3-methyl-2-butenyl)amino]methyl]amino]butyl]- (9CI) (CA INDEX NAME)

IT 365568-09-8P

RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); PROC (Process); RACT (Reactant or reagent)

(design of antihypertensive drugs from Verbesina caracasana)

RN 365568-09-8 CAPLUS

CN 2-Propenamide, N-[6-[(aminoiminomethyl)amino]hexyl]-3-(3,4-dimethoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

MeO
$$E$$
 $(CH_2)_6$ N NH_2 NH_2

IT 14737-89-4

RL: RCT (Reactant); RACT (Reactant or reagent)

(design of antihypertensive drugs from Verbesina caracasana)

RN 14737-89-4 CAPLUS

CN 2-Propenoic acid, 3-(3,4-dimethoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

REFERENCE COUNT: 32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 3 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1997:752957 CAPLUS

DOCUMENT NUMBER: 128:34753

TITLE: Preparation of antihypertensive carboline

derivatives

INVENTOR(S): Bombrun, Agnes

PATENT ASSIGNEE(S): Icos Corporation, USA; Bombrun, Agnes

SOURCE: PCT Int. Appl., 122 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

| PA. | FENT | NO. | | | KIN | D - | DATE | | | APPL: | ICAT | ION I | . 00 | | D | ATE | |
|-----|------|-----|-----|-----|-----------|----------|------|----------------|-----|-------|-------|-------|------|-----|-------|-------|-----|
| WO | 9743 | 287 | | | A1 | 19971120 | | WO 1997-EP2277 | | | | | | 1: | 9970: | 505 | |
| | W: | AL, | AM, | ΑT, | ΑU, | ΑZ, | BA, | BB, | BG, | BR, | BY, | CA, | CH, | CN, | CU, | CZ, | DE, |
| | | DK, | EE, | ES, | FI, | GB, | GE, | GH, | HU, | IL, | IS, | JP, | KΕ, | KG, | ΚP, | KR, | ΚZ, |
| | | LC, | LK, | LR, | LS, | LT, | LU, | LV, | MD, | MG, | MK, | MN, | MW, | MX, | NO, | NZ, | PL, |
| | | PT, | RO, | RU, | SD, | SE, | SG, | ·SI, | SK, | ТJ, | TM, | TR, | TT, | UA, | UG, | US, | UZ, |
| | | VN, | ΥU | | | | | | | | | | | | | | |
| | RW: | GH, | KΕ, | LS, | MW, | SD, | SZ, | UG, | ΑT, | BE, | CH, | ĎΕ, | DK, | ES, | FI, | FR, | GB, |
| | | GR, | ΙE, | IT, | LU, | MC, | NL, | PT, | SE, | BF, | ВJ, | CF, | CG, | CI, | CM, | GA, | GN, |
| | | ML, | MR, | ΝE, | SN, | TD, | TG | | | | | | | | | | |
| CA | 2253 | 948 | | | A1 | | 1997 | 1120 | (| CA 19 | 997-: | 2253 | 948 | | 19 | 9970 | 505 |
| CA | 2253 | 948 | | | С | | 2005 | 0726 | | | | | | | | | |
| ΑU | 9728 | 910 | | | Α | | 1997 | 1205 | 7 | AU 19 | 997-2 | 2891 |) | | 19 | 9970! | 505 |
| AU | 7118 | 85 | | | B2 | | 1999 | 1021 | | | | | | | | | |
| EΡ | 9125 | 67 | | | A1 | | 1999 | 0506 |] | EP 19 | 997- | 92296 | 50 | | 1.5 | 9970 | 505 |
| EP | 9125 | 67 | | | B1 | | 2002 | 0410 | | | | | | | | | |
| | R: | ΑT, | BE, | CH, | DE, | DK, | ES, | FR, | GB, | GR, | IT, | LI, | LU, | NL, | SE, | MC, | PT, |
| | | | - | | LV, | | | | | | | | | | | | |
| | 1218 | | | | | | | | (| CN 19 | 997-: | 19450 | 80 | | 19 | 9970 | 505 |
| CN | 1067 | 071 | | | В | | 2001 | 0613 | | | | | | | | | |

| BR 9709230 | Α | 19990810 | BR | 1997-9230 | | 19970505 |
|------------------------|---------|----------|----|-------------|----|----------|
| HU 9901478 | A2 | 19990830 | HU | 1999-1478 | | 19970505 |
| JP 2000513717 | ${f T}$ | 20001017 | JP | 1997-540456 | | 19970505 |
| JP 3418405 | B2 | 20030623 | | | | |
| MD 980248 | Α | 20001031 | MD | 1998-248 | | 19970505 |
| AT 215950 | ${f T}$ | 20020415 | ΑT | 1997-922960 | | 19970505 |
| ES 2175404 | T3 | 20021116 | ES | 1997-922960 | | 19970505 |
| US 6043252 | Α | 20000328 | US | 1998-154052 | | 19980916 |
| NO 9805222 | Α | 19990111 | NO | 1998-5222 | | 19981109 |
| KR 2000010918 | Α | 20000225 | KR | 1998-709066 | | 19981110 |
| US 6117881 | A | 20000912 | US | 1999-155811 | | 19990423 |
| US 6306870 | B1 | 20011023 | US | 2000-592514 | | 20000612 |
| PRIORITY APPLN. INFO.: | | | GB | 1996-9777 | Α | 19960510 |
| | | • | GB | 1996-9820 | Α | 19960510 |
| | | • | WO | 1997-EP2277 | W | 19970505 |
| | | | US | 1999-155811 | A2 | 19990423 |

OTHER SOURCE(S):

MARPAT 128:34753

I

GΙ

$$\begin{array}{c|c}
R & & & \\
& & & \\
N & & & \\
H & & & \\
R^2 & O & & \\
\end{array}$$

Carboline derivs. I [R = H, halogen; R1, R2 = (un) substituted Ph] were prepared and are potent and selective inhibitors of cyclic guanosine 3',5'-monophosphate specific phosphodiesterase (cGMP specific PDE). Thus, tryptamine was cyclized with piperonal and treated with (E)-HO2CCH:CHC6H4NHAc-4 to give I [R = H, R1 = 4-AcNHC6H4, R2 = 3,4-methylenedioxyphenyl, II]. II had an IC50 for cGMP-PDE inhibition of 5 nM.

IT 501-16-6, (E)-3,4-Dihydroxycinnamic acid 537-98-4

11 501-16-6, (E)-3,4-Dinydroxycinnamic acid 537-98-4 14737-89-4, (E)-3,4-Dimethoxycinnamic acid 20329-98-0 RL: RCT (Reactant); RACT (Reactant or reagent) (preparation of antihypertensive carboline derivs.) RN 501-16-6 CAPLUS

CN 2-Propenoic acid, 3-(3,4-dihydroxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 537-98-4 CAPLUS

CN 2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

RN 14737-89-4 CAPLUS

CN 2-Propenoic acid, 3-(3,4-dimethoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 20329-98-0 CAPLUS

CN 2-Propenoic acid, 3-(3,4,5-trimethoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

IT 86981-09-1P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of antihypertensive carboline derivs.)

RN 86981-09-1 CAPLUS

CN 2-Propenoic acid, 3-(4-hydroxy-3-methoxy-5-nitrophenyl)-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L9 ANSWER 4 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1992:83664 CAPLUS

DOCUMENT NUMBER:

116:83664

TITLE:

Preparation of 5,6,7,8-tetrahydro-4H-thiazolo[5,4-

b]azepine derivatives as antihypertensives

INVENTOR(S):

Aono, Tetsuya; Shimamoto, Norio

PATENT ASSIGNEE(S):

Takeda Chemical Industries, Ltd., Japan

SOURCE:

Jpn. Kokai Tokkyo Koho, 63 pp.

CODEN: JKXXAF

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT:

1

FAMILI ACC. NOM. COC

PATENT INFORMATION:

| PATENT NO. | ĶIND | DATE | APPLICATION NO. | DATE |
|------------------------|-------------|-----------|-----------------|----------|
| | - - | | | |
| JP 03206042 | Α | 19910909 | JP 1990-833 | 19900106 |
| PRIORITY APPLN. INFO.: | | | JP 1990-833 | 19900106 |
| OTHER SOURCE(S): | MARPAT | 116:83664 | | |
| GI | | | | |

Ι

AB The title compds. [I; R1 = H, (un)substituted aliphatic, acyl or sulfonyl; R2 = H, (un)substituted aromatic or aliphatic] are prepared as K channel opener. Thus, 14.8 g 1,1'-carbonyldiimidazole was added to a solution of 12 g 2,6-F2C6H3CO2H in THF and thereto after stirring 15 min at room temperature 9.73

g 3-amino- ϵ -caprolactam was added and the mixture was stirred 5 h at room temperature to give 13.5 g 3-(2,6-difluorobenzoylamino)'- ϵ -caprolactam which (8.96 g) was refluxed 24 h, with 8.96 g P4S10 in pyridine to give 23.8% I (R1 = H, R2 = 2,6-F2C6H3)(II). II and I [R1 = H, R2 = (Z)-4-Et2NC6H4CH:CH] (III) in vitro inhibited 8 and 100%, resp., rat aorta contraction induced by Et3NCl and BaCl2 and gave no inhibition of the one induced by 80 mM KCl. II and III at 1 mg/kg i.v. lowered 49 and 46%, resp. the blood pressure of rats. A total of 175 I were prepared 537-73-5, 3-Hydroxy-4-methoxycinnamic acid 128069-93-2

RL: RCT (Reactant); RACT (Reactant or reagent)

(acylation by, of aminocaprolactam)

RN 537-73-5 CAPLUS

IT

CN 2-Propenoic acid, 3-(3-hydroxy-4-methoxyphenyl)- (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{CH-CO}_2\text{H} \\ \\ \text{OH} \end{array}$$

RN 128069-93-2 CAPLUS

CN 2-Propenoic acid, 3-[3-methoxy-4-[(methylthio)methoxy]phenyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{OMe} \\ \text{MeS-CH}_2\text{-O} \\ \text{CH-CH-CO}_2\text{H} \end{array}$$

IT 128068-07-5P 128068-08-6P 128068-15-5P

128068-16-6P

RL: SPN (Synthetic preparation); PREP (Preparation)

(preparation and sulfuration-cyclization of, antihypertensive

tetrahydrothiazoloazepine derivative from)

RN 128068-07-5 CAPLUS

CN 2-Propenamide, N-(hexahydro-2-oxo-1H-azepin-3-yl)-3-(2,3,4-

trimethoxyphenyl) - (9CI) (CA INDEX NAME)

RN 128068-08-6 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-(hexahydro-2-oxo-1H-azepin-3-yl)-

(9CI) (CA INDEX NAME)

RN 128068-15-5 CAPLUS

CN 2-Propenamide, N-(hexahydro-2-oxo-1H-azepin-3-yl)-3-[4-methoxy-3-[2-(methylthio)ethoxy]phenyl]- (9CI) (CA INDEX NAME)

RN 128068-16-6 CAPLUS

CN 2-Propenamide, N-(hexahydro-2-oxo-1H-azepin-3-yl)-3-(3,4,5-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

L9 ANSWER 5 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1983:83222 CAPLUS

DOCUMENT NUMBER:

98:83222

TITLE:

Pyrimidine derivatives. 4. Synthesis and

antihypertensive activity of

4-amino-2-(4-cinnamoylpiperazino)-6,7-

dimethoxyquinazoline derivatives

Sekiya, Tetsuo; Hiranuma, Hidetoshi; Hata, Shunsuke; AUTHOR (S):

Mizogami, Susumu; Hanazuka, Mitsuo; Yamada, Shunichi Res. Lab., Mitsubishi Yuka Pharmaceutical Co., Ltd.,

CORPORATE SOURCE: Ibaraki, 300-03, Japan

Journal of Medicinal Chemistry (1983), 26(3), 411-16 SOURCE:

CODEN: JMCMAR; ISSN: 0022-2623

Journal DOCUMENT TYPE: English LANGUAGE:

OTHER SOURCE(S): CASREACT 98:83222

GT

AB The title compds. I (R = H or Me; R1 = Ph, substituted Ph, furyl or thienyl) as the HCl salts, prepared either by condensation of 4-amino-2-chloro-6,7-dimethoxyquinazoline [23680-84-4] with acryloylpiperazines or by selective acylation of 4-amino-6,7-dimethoxy-2piperazinoquinazoline-HCl [84050-22-6] with mixed anhydrides, were evaluated for their ability to reduce blood pressure in conscious, spontaneously hypertensive rats. 4-Amino-2-(4cinnamoylpiperazino)-6,7-dimethoxyquinozoline [70842-66-9] Showed activity at oral doses 0.3-10 mg/kg in the above rats, and at 3 and 10 mg in renal hypertensive rats, and α -adrenoceptor blocking effects in isolated aorta of rats. Structure-activity relations are discussed.

Ι

33130-03-9 IT

> RL: RCT (Reactant); RACT (Reactant or reagent) (reaction of, with piperizine hydrobromide)

RN 33130-03-9 CAPLUS

2-Propenoic acid, 3-(2,3,4-trimethoxyphenyl)- (9CI) CN (CA INDEX NAME)

ANSWER 6 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2004:995933 CAPLUS

DOCUMENT NUMBER: 141:424343

TITLE: Preparation of nitrosated and nitrosylated compounds

for use in pharmaceutical compositions a nitric oxide

(NO) donors

INVENTOR (S): Earl, Richard A.; Garvey, David S.; Gaston, Ricky D.;

Lin, Chia-En; Ranatunge, Ramani R.; Richardson,

Stewart K.; Stevenson, Cheri A.

PATENT ASSIGNEE(S):

Nitromed, Inc., USA

SOURCE: PCT Int. Appl., 181 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | KIND DATE | APPLICATION NO. | DATE | | | |
|------------------------|-----------------|---------------------|-----------------|--|--|--|
| | A2 20041118 | WO 2004-US7943 | 20040315 | | | |
| W: AE, AG, AL, | AM, AT, AU, AZ, | BA, BB, BG, BR, BW, | BY, BZ, CA, CH, | | | |
| CN, CO, CR, | CU, CZ, DE, DK, | DM, DZ, EC, EE, EG, | ES, FI, GB, GD, | | | |
| | | IN, IS, JP, KE, KG, | | | | |
| LK, LR, LS, | LT, LU, LV, MA, | MD, MG, MK, MN, MW, | MX, MZ, NA, NI, | | | |
| | | RO, RU, SC, SD, SE, | | | | |
| | | UG, US, UZ, VC, VN, | | | | |
| RW: BW, GH, GM, | KE, LS, MW, MZ, | SD, SL, SZ, TZ, UG, | ZM, ZW, AM, AZ, | | | |
| BY, KG, KZ, | MD, RU, TJ, TM, | AT, BE, BG, CH, CY, | CZ, DE, DK, EE, | | | |
| · ES, FI, FR, | GB, GR, HU, IE, | IT, LU, MC, NL, PL, | PT, RO, SE, SI, | | | |
| SK, TR, BF, | BJ, CF, CG, CI, | CM, GA, GN, GQ, GW, | ML, MR, NE, SN, | | | |
| TD, TG | | | • | | | |
| AU 2004237574 | A1 20041118 | AU 2004-237574 | 20040315 | | | |
| CA 2518506 | A1 20041118 | CA 2004-2518506 | 20040315 | | | |
| EP 1603933 | A2 20051214 | EP 2004-749385 | 20040315 | | | |
| R: AT, BE, CH, | DE, DK, ES, FR, | GB, GR, IT, LI, LU, | NL, SE, MC, PT, | | | |
| IE, SI, LT, | LV, FI, RO, MK, | CY, AL, TR, BG, CZ, | EE, HU, PL, SK | | | |
| US 2006009431 | A1 20060112 | US 2005-221901 | 20050909 | | | |
| PRIORITY APPLN. INFO.: | | US 2003-453963P | P 20030313 | | | |
| | | US 2003-482134P | P 20030625 | | | |
| | | WO 2004-US7943 | A 20040315 | | | |
| OTHER SOURCE(S): | MARPAT 141:4243 | 43 | | | | |

GI

AΒ Nitroso and nitrosyl derivs. of therapeutic agents, such as R-SNO, R-ONO, R-ONO2 [R = antithrombogenic agent, thrombolytic agent, fibrinolytic agent, vasospasm inhibitor, potassium channel blocker, calcium channel blocker, antihypertensive agent, antimicrobial agent, antibiotic, platelet reducing agent, antimitotic agent, antiproliferative agent, microtubule inhibitor, antisecretory agent, remodeling inhibitor, antisense nucleotide, anticancer chemotherapeutic agent, steroid, nonsteroidal antiinflammatory agent, selective COX-2 inhibitor, immunosuppressive agent, growth factor antagonist or antibody, dopamine agonist, radiotherapeutic agent, heavy metal functioning as a radioplaque agent, biol. agent, aldosterone antagonist, α -adrenergic receptor antagonist, angiotensin II antagonist, β-adrenergic agonist, antihyperlipidemic drug, angiotensin converting enzyme (ACE) inhibitor, antioxidant, \(\beta\)-adrenergic antagonist, endothelin antagonist, neutral endopeptidase inhibitor, renin inhibitor, free radical scavenger, iron chelator, sex hormone, antipolymerase, antiviral agent, photodynamic therapy agent, antibody targeted therapy agent, gene therapy agent, etc.], were prepared for therapeutic use. The compds. and compns. of this

invention can also be bound to a matrix. These nitroso- and nitro-compds. are claimed for use in treating cardiovascular diseases, for inhibiting platelet aggregation and platelet adhesion caused by the exposure of blood to a medical device, for treating pathol. conditions resulting from abnormal cell proliferation; transplantation rejections, autoimmune, inflammatory, proliferative, hyperproliferative or vascular diseases; for reducing scar tissue or for inhibiting wound contraction, particularly the prophylactic and/or therapeutic treatment of restenosis by administering at least one compound of the invention that is optionally nitrosated and/or nitrosylated, in combination with nitric oxide donors that are capable of releasing nitric oxide or indirectly delivering or transferring nitric oxide to targeted sites under physiol. conditions. The compds. of this invention are preferably estradiol compds., troglitazone compds., tranilast compds., retinoic acid compds., resveratrol compds., mycophenolic acid compds., acid compds., anthracenone compds. and trapidil compds. The cardiovascular diseases for treatment include restenosis, coronary artery disease, atherosclerosis, atherogenesis, cerebrovascular disease, angina, ischemic disease, congestive heart failure or pulmonary edema associated with acute myocardial infarction, aneurysm, thrombosis, hypertension, platelet adhesion, platelet aggregation, smooth muscle cell proliferation, a vascular or non-vascular complication associated with the use of a medical device, wounds associated with the use of a medical device, pulmonary thromboembolism, cerebral thromboembolism, thrombophlebitis, thrombocytopenia or a bleeding disorder. The autoimmune diseases for treatment include a pathol. condition resulting from abnormal cell proliferation, polycystic kidney disease, an inflammatory disease, for preserving an organ and/or a tissue or for inhibiting wound contraction in a patient. The pathol. conditions resulting from abnormal cell proliferation include is a cancer, a Karposi's sarcoma, a cholangiocarcinoma, a choriocarcinoma, a neoblastoma, a Wilm's tumor, Hodgkin's disease, a melanoma, multiple myelomas, a chronic lymphocytic leukemia or an acute or chronic granulocytic lymphoma. The inflammatory diseases for treatment includerheumatoid arthritis, an inflammatory skin disease, restenosis, multiple sclerosis, a surgical adhesion, tuberculosis, a graft rejection, an inflammatory lung disease, an inflammatory bowel disease, an inflammatory disease that affects or causes obstruction of a body passageway, an inflammation of the eye, an inflammation of the nose, an inflammation of the throat or a neovascular diseases of the eye. Thus, S-mono- and O,S-dinitroso- β -estradiol derivs. I (R = NO, R1 = H, NO) were prepared via an esterification reaction of β -estradiol with 3-methyl-3-(2,4,6-trimethoxyphenylmethylthio) buty ric acid using EDAP and DMAP in DMF to form mono-ester I [R = CH2C6H2-2,4,6-(OMe)3, R1 = H], cleavage of the trimethoxybenzyl S-protecting group of the mono-ester using L-cysteine and TFA in CH2Cl2 to give thiol I (R = R1 = H), and finally, treatment of the thiol with Bu nitrite in CH2Cl2 to form the desired S-mono- and O,S-dinitroso- β estradiol derivs. The prepared compds. were assayed for suppression of proliferation of human coronary artery smooth muscle cells. 53902-12-8DP, Tranilast, derivs. 794519-34-9P 794519-35-0P 794519-36-1P 794519-37-2P RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(preparation of nitrosated and nitrosylated compds. for use in pharmaceutical compns. as nitric oxide (NO) donors) 53902-12-8 CAPLUS

IT

RN

Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

RN 794519-34-9 CAPLUS

CN Benzoic acid, 2-[[(2E)-3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]-, 2-[[2-methyl-2-(nitrosothio)propyl]amino]-2-oxoethyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 794519-35-0 CAPLUS

CN Benzoic acid, 2-[[(2E)-3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]-, 2-[3-methyl-3-(nitrosothio)butoxy]-2-oxoethyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 794519-36-1 CAPLUS

CN Benzoic acid, 2-[[(2E)-3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]-, 2-[4-[2-methyl-2-(nitrosothio)propyl]-1-piperazinyl]-2-oxoethyl ester (9CI) (CA INDEX NAME)

RN 794519-37-2 CAPLUS

CN Benzoic acid, 2-[[(2E)-3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]-, 2-[2-[4-[2-methyl-2-(nitrosothio)propyl]-1-piperazinyl]ethoxy]-2-oxoethyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

PAGE 1-B

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__NO
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RN 794519-94-1 CAPLUS

CN Benzoic acid, 2-[[(2E)-3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]-, 2-(1,1-dimethylethoxy)-2-oxoethyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 794519-95-2 CAPLUS

CN Benzoic acid, 2-[[(2E)-3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]-, carboxymethyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

$$HO_2C$$
 OMe

RN 794519-96-3 CAPLUS

CN Benzoic acid, 2-[[(2E)-3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]-, 2-[(2-mercapto-2-methylpropyl)amino]-2-oxoethyl ester (9CI) (CA INDEX NAME)

RN 794519-97-4 CAPLUS

CN Benzoic acid, 2-[[(2E)-3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]-, 2-(3-mercapto-3-methylbutoxy)-2-oxoethyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 794519-98-5 CAPLUS

CN Benzoic acid, 2-[[(2E)-3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]-, 2-[4-(2-mercapto-2-methylpropyl)-1-piperazinyl]-2-oxoethyl ester (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L9 ANSWER 7 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2001:152482 CAPLUS

DOCUMENT NUMBER: 134:157568

TITLE: Agent inhibiting hypertensive arteriolar

disorder

INVENTOR(S): Iwaki, Yoichi; Kusama, Hiroshi; Tsuji, Atsutoshi

PATENT ASSIGNEE(S): Kissei Pharmaceutical Co., Ltd., Japan

SOURCE: PCT Int. Appl., 11 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

DATE APPLICATION NO. DATE PATENT NO. KIND -----_ _ _ _ -----______ _ _ _ _ _ _ _ _ 20010301 WO 2000-JP4528 20000707 WO 2001013911 A1

W: JP, US

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE

PRIORITY APPLN. INFO.:

JP 1999-233008

This document discloses an agent inhibiting diseases concerning hypertensive arteriolar disorder (cerebral stroke, vascular dementia, hypertensive eyeground, hypertensive retinopathy, etc.) containing as the active ingredient N-(3,4dimethoxycinnamoyl)anthranilic acid (tranilast), which has effects of remarkably inhibiting arteriolar basement membrane thickening caused by hypertension etc., or pharmacol. acceptable salts thereof.

53902-12-8, Tranilast TΤ

> RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(agent inhibiting hypertensive arteriolar disorder)

53902-12-8 CAPLUS RN

Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) CN (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 8 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

8

ACCESSION NUMBER:

2005:1242303 CAPLUS

DOCUMENT NUMBER:

143:477660

TITLE:

Preparation of cyclohexyldiamine derivatives as

modulators of ORL1 receptors

INVENTOR(S):

Sundermann, Corinna; Sundermann, Bernd

PATENT ASSIGNEE(S): Gruenenthal G.m.b.H., Germany

SOURCE:

PCT Int. Appl., 74 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO | D. DATE |
|---------------|--------------|-----------|-------------------|-----------------------|
| | - | | | · |
| WO 2005110975 | A1 | 20051124 | WO 2005-EP4912 | 2 20050506 |
| W: AE, AG, | AL, AM, AT | , AU, AZ, | BA, BB, BG, BR, B | BW, BY, BZ, CA, CH, |
| CN, CO, | CR, CU, CZ | , DK, DM, | DZ, EC, EE, EG, E | ES, FI, GB, GD, GE, |
| GH, GM, | HR, HU, ID | , IL, IN, | IS, JP, KE, KG, H | KM, KP, KR, KZ, LC, |
| LK, LR, | LS, LT, LU | , LV, MA, | MD, MG, MK, MN, N | NW, MX, MZ, NA, NI, |
| NO, NZ, | OM, PG, PH | , PL, PT, | RO, RU, SC, SD, S | SE, SG, SK, SL, SM, |
| SY, TJ, | TM, TN, TR | , TT, TZ, | UA, UG, US, UZ, V | C, VN, YU, ZA, ZM, ZW |
| RW: BW, GH, | GM, KE, LS | , MW, MZ, | NA, SD, SL, SZ, T | TZ, UG, ZM, ZW, AM, |
| | | | | CH, CY, CZ, DE, DK, |
| EE, ES, | FI, FR, GB | , GR, HU, | IE, IS, IT, LT, I | LU, MC, NL, PL, PT, |
| | | | | SA, GN, GQ, GW, ML, |
| | SN, TD, TG | | | |

DE 2004-102004023506 DE 102004023506 **A**1 20051201 20040510 CA 2566219 CA 2005-2566219 Δ1 20051124 20050506 EP 2005-747800 EP 1747191 A1 20070131 20050506 AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, LV PRIORITY APPLN. INFO.: DE 2004-102004023506A 20040510 WO 2005-EP4912 W 20050506

OTHER SOURCE(S): MARPAT 143:477660

GI

RN

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Title compds. I [n = 1-2; R1 and R2 independently = H, (un) substitutedalkyl, cycloalkyl, etc. or R1 and R2 together may form CH2CH2OCH2CH2, CH2CH2NR5CH2CH2 or (CH2)3-6; R5 = H, (un)substituted alkyl, aryl, etc.; R3 = (un)substituted alkyl, cycloalkyl, heteroaryl, etc.; R4 = (un) substituted alkyl, heteroaryl, aryl, etc.; X = (CR6R7) m; m = 0-4; A = NH, O, S, etc.; R6 and R7 independently = H, (un)substituted alkyl or aryl with provisions] and their pharmaceutically acceptable salts, are prepared and disclosed as modulators of ORL1 receptors. Thus, e.g., II was prepared by coupling of N, N-dimethyl-1-phenylcyclohexan-1, 4-diamine with 4-phenoxybutyrylchloride and subsequent conversion into the hydrochloride. The binding activity of I towards ORL1 receptors was evaluated in scintillation assays using recombinant CHO-ORL1 cells and it was revealed that selected compds. of the invention displayed binding activity in the range of 43 up to 99%. I as modulator of ORL1 receptors should prove useful in the treatment of obesity, depression and pain. Pharmaceutical compns. comprising I are disclosed.

IT 869798-66-3P 869798-69-6P 869798-83-4P 869798-84-5P 869798-86-7P 869799-14-4P 869799-18-8P 869799-20-2P 869799-21-3P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of cyclohexyldiamine derivs. as modulators of ORL1 receptors) 869798-66-3 CAPLUS

2 - Propenamide, 3 - [3 - methoxy - 4 - (phenylmethoxy) phenyl] - N - [4 - phenyl - 4 - (1 - piperidinyl) cyclohexyl] - (9CI) (CA INDEX NAME)

RN 869798-69-6 CAPLUS

CN 2-Propenamide, N-[4-(dimethylamino)-4-(phenylmethyl)cyclohexyl]-3-[3-methoxy-4-(phenylmethoxy)phenyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} \text{CH} & \text{CH} & \text{CH} \\ \text{CH} & \text{CH} & \text{CH} \\ \text{OMe} \end{array}$$

RN 869798-83-4 CAPLUS

CN 2-Propenamide, N-[4-(dimethylamino)-4-[(3-methylphenyl)methyl]cyclohexyl]-3-[3-methoxy-4-(phenylmethoxy)phenyl]- (9CI) (CA INDEX NAME)

RN 869798-84-5 CAPLUS

CN 2-Propenamide, N-[4-(dimethylamino)-4-[(4-methylphenyl)methyl]cyclohexyl]-3-[3-methoxy-4-(phenylmethoxy)phenyl]- (9CI) (CA INDEX NAME)

Me
$$O = CH_2 - Ph$$
 $O = CH_2 - Ph$
 $O = CH_2 - Ph$
 $O = CH_2 - Ph$
 $O = CH_2 - Ph$

RN 869798-86-7 CAPLUS

CN 2-Propenamide, N-[4-[(3-chlorophenyl)methyl]-4-(dimethylamino)cyclohexyl]-3-[3-methoxy-4-(phenylmethoxy)phenyl]- (9CI) (CA INDEX NAME)

RN 869799-14-4 CAPLUS

CN 2-Propenamide, N-[4-(dimethylamino)-4-[(3-fluorophenyl)methyl]cyclohexyl]-3-[3-methoxy-4-(phenylmethoxy)phenyl]- (9CI) (CA INDEX NAME)

RN 869799-18-8 CAPLUS

CN 2-Propenamide, N-[4-(dimethylamino)-4-(2-phenylethyl)cyclohexyl]-3-[3-methoxy-4-(phenylmethoxy)phenyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} \text{CH} & \text{CH} & \text{CH} & \text{CH} \\ \hline \\ \text{Ph} & \text{CH}_2 - \text{O} \\ \hline \\ \text{OMe} \\ \end{array}$$

RN 869799-20-2 CAPLUS

CN 2-Propenamide, N-[4-[(4-chlorophenyl)methyl]-4-(dimethylamino)cyclohexyl]-3-[3-methoxy-4-(phenylmethoxy)phenyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} \text{OMe} & \text{OMe} \\ \text{O} & \text{O-CH}_2\text{-Ph} \\ \text{CH}_2 & \text{NMe}_2 \end{array}$$

RN 869799-21-3 CAPLUS

CN 2-Propenamide, N-[4-(dimethylamino)-4-[(2-fluorophenyl)methyl]cyclohexyl]-3-[3-methoxy-4-(phenylmethoxy)phenyl]- (9CI) (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 9 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

6

ACCESSION NUMBER: 2004:514592 CAPLUS

DOCUMENT NUMBER:

141:17191

TITLE:

Tranilast attenuates myocardial fibrosis in

association with suppression of monocyte/macrophage

infiltration in DOCA/salt hypertensive rats

Kagitani, Satoshi; Ueno, Hitoshi; Hirade, Satoshi;

Takahashi, Toru; Takata, Masanobu; Inoue, Hiroshi Second Department of Internal Medicine, Toyama Medical

and Pharmaceutical University, Toyama, Japan

Journal of Hypertension (2004), 22(5), 1007-1015

CODEN: JOHYD3; ISSN: 0263-6352

PUBLISHER: Lippincott Williams & Wilkins

DOCUMENT TYPE: Journal LANGUAGE: English

AUTHOR (S):

SOURCE:

CORPORATE SOURCE:

Objective In order to study the association between myocardial fibrosis and ΔR inflammatory cell infiltration in the hypertensive heart, we investigated whether N(3,4-dimethoxycinnamoyl) anthranilic acid (tranilast), an anti-inflammatory drug, would suppress myocardial fibrosis via inhibition of inflammatory cell infiltration in deoxycorticosteroneacetate (DOCA) hypertensive rats. Methods Sprague-Dawley rats treated with DOCA combined with the addition of 1% NaCl and 0.2% KCl in the drinking water after left nephrectomy were given tranilast (100 mg/kg per day, n = 15) or vehicle (n = 15) for up to 4 wk. Systolic blood pressure (SBP), amount of myocardial interstitial fibrosis, perivascular fibrosis and type I and III collagen, and mRNA expression of procollagen I (PI) and procollagen III (PIII), transforming growth factor (TGF)-β1, type-1 plasminogen activator inhibitor (PAI-1), monocyte chemoattractant protein (MCP)-1 and interleukin (IL)-6 were determined Results SBP was increased significantly 2 wk after treatment with DOCA and salt. Myocardial interstitial fibrosis, perivascular fibrosis and collagen accumulation increased significantly 4 wk after the treatment. Two weeks after the treatment with DOCA and salt, mRNA expression of PI and PIII, TGF- β 1, PAI-1, MCP-1 and IL-6 increased significantly. Although the SBP was similar in animals treated with tranilast or vehicle, monocyte/macrophage infiltration was suppressed, mRNA expression of TGF-β1, PAI-1, MCP-1, IL-6, PI and PIII was attenuated, and myocardial fibrosis and collagen accumulation were suppressed in hypertensive animals receiving tranilast. Conclusion Myocardial fibrosis seen in DOCA/salt hypertensive rats might be associated with the inflammation/wound healing response. Tranilast suppresses both infiltration of monocytes/macrophages and myocardial fibrosis.

IT 53902-12-8, Tranilast

RL: DMA (Drug mechanism of action); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(tranilast attenuates myocardial fibrosis in association with suppression of monocyte/macrophage infiltration in DOCA/salt hypertensive rats)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

REFERENCE COUNT:

31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 10 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1993:625746 CAPLUS

DOCUMENT NUMBER: 119:225746

TITLE:

Novel hypotensive agents from Verbesina caracasana. 2.

Synthesis and pharmacology of caracasanamide

AUTHOR (S):

Delle Monache, Giuliano; Botta, Bruno; Delle Monache, Franco; Espinal, Romulo; De Bonnevaux, Stella C.; De Luca, Carlo; Botta, Maurizio; Corelli, Federico;

Carmignani, Marco

CORPORATE SOURCE:

SOURCE:

Cent. Chim. Recett., Rome, 00168, Italy

Journal of Medicinal Chemistry (1993), 36(20), 2956-63

CODEN: JMCMAR; ISSN: 0022-2623

DOCUMENT TYPE:

LANGUAGE:

Journal English

GI

ΔR Caracasanamide, one of the hypotensive agents isolated from Verbesiana caracasana, is a mixture of (Z) - and (E) -I. The structure of (E) -I was confirmed by high-yielding synthesis starting from N,N'-bis(tertbutoxycarbonyl)-S-methylisothiourea. The water-soluble (Z)-I, assayed at 50 to 1600 µg/kg i.v. in rats, decreased blood pressure, increased cardiac inotropism, respiratory frequency, and tidal volume, and induced a very slight, insignificant tachycardia. Higher doses produced respiratory depression and, in some cases, consequent cardiac arrest. (Z)-I affects cardiovascular function by acting at the vascular level in inducing arterial vasodilation, by determining sympathetic hypotone through central neurogenic mechanisms, and by interacting with the cardiac β 1-adrenoreceptors. The respiratory effects were independent of the cardiovascular ones. In lowering blood pressure, (Z)-I was more potent than guanethidine and no less potent than reserpine and papaverine. may therefore be useful in the treatment of arterial hypertension of moderate degree.

IT 14737-88-3P, (Z)-3,4-Dimethoxycinnamic acid 14737-89-4P,
 (E)-3,4-Dimethoxycinnamic acid 150785-42-5P 150785-43-6P
 RL: FORM (Formation, nonpreparative); PREP (Preparation)
 (formation of, from caracasanamide)

RN 14737-88-3 CAPLUS

CN 2-Propenoic acid, 3-(3,4-dimethoxyphenyl)-, (2Z)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 14737-89-4 CAPLUS

CN 2-Propenoic acid, 3-(3,4-dimethoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

RN 150785-42-5 CAPLUS

CN 2-Propenamide, N-[4-[(aminocarbonyl)amino]butyl]-3-(3,4-dimethoxyphenyl)-, (Z)-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

$$\begin{array}{c|c} & H & H \\ \hline Z & O & \\ \hline \\ O & O \\ \hline \\ O & O \\ \end{array}$$

RN 150785-43-6 CAPLUS

CN 2-Propenamide, N-[4-[(aminocarbonyl)amino]butyl]-3-(3,4-dimethoxyphenyl)-, (E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

MeO
$$E$$
 N
 H
 $CH_2)_4$
 N
 H
 NH_2

IT 150785-46-9P

RL: SPN (Synthetic preparation); PREP (Preparation) (intermediate in preparation of caracasanamide)

RN 150785-46-9 CAPLUS

CN Carbamic acid, [[[4-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]butyl]amino][[(1,1-dimethylethoxy)carbonyl]imino]methyl](3-methyl-2-butenyl)-, 1,1-dimethylethyl ester, (E)- (9CI) (CA INDEX NAME)

IT 146269-39-8P, (E)-Caracasanamide

RL: PREP (Preparation)

(isolation and mol. structure of)

RN 146269-39-8 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[[imino[(3-methyl-2-butenyl)amino]methyl]amino]butyl]-, (E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

IT 146269-40-1P, (Z)-Caracasanamide

RL: PREP (Preparation)

(isolation, mol. structure, and antihypertensive activity of)

RN 146269-40-1 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[[imino[(3-methyl-2-butenyl)amino]methyl]amino]butyl]-, (2Z)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

$$\begin{array}{c|c} & H & H & H \\ \hline Z & O & NH & N \\ \hline \\ O & OMe & NH & N \\ \hline \end{array}$$

L9 ANSWER 11 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:523280 CAPLUS

DOCUMENT NUMBER: 143:59817

TITLE: Preparation of nitrooxy derivatives of carvedilol and

other β -blockers as antihypertensive

drugs

INVENTOR(S): Del Soldato, Piero; Benedini, Francesca; Ongini, Ennio

PATENT ASSIGNEE(S): Nicox S. A., Fr.

SOURCE: PCT Int. Appl., 103 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

LANGUAGE: Engli: FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PAT | ENT | NO. | | | KIN | D : | DATE | | 2 | APPL | ICAT | ION | NO. | | D | ATE | |
|-----|------|--------------|-----|-----|-----|------------|------|------|-----|------|------|------|-----|-----|-----|------|-----|
| | | - | | | | _ | | | | | | | | | - | | |
| WO | 2005 | 0536 | 85 | | A1 | : | 2005 | 0616 | 1 | WO 2 | 004- | EP13 | 683 | | 2 | 0041 | 201 |
| | W: | ΑE, | AG, | AL, | AM, | ΑT, | AU, | ΑZ, | BA, | BB, | BG, | BR, | BW, | BY, | BZ, | CA, | CH, |
| | | | | | CU, | | | | | | | | | | | | |
| | | GE, | GH, | GM, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | KΕ, | KG, | KΡ, | KR, | ΚZ, | LC, |
| | | LK, | LR, | LS, | LT, | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | MZ, | NA, | NI, |
| | | NO, | NZ, | OM, | PG, | PH, | PL, | PT, | RO, | RU, | SC, | SD, | SE, | SG, | SK, | SL, | SY, |
| | | | | | TR, | | | | | | | | | | | | |

```
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
             EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT,
             RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,
             MR, NE, SN, TD, TG
                                             AU 2004-294297
                                20050616
     AU 2004294297
                          A1
                                                                    20041201
     CA 2548129
                          A1
                                20050616
                                             CA 2004-2548129
                                                                    20041201
                                20060823
                                             EP 2004-803434
     EP 1691804
                          A1
                                                                    20041201
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
         R:
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK,
             BA, HR, IS, YU
     CN 1886132
                                20061227
                                             CN 2004-80035459
                                                                    20041201
                          Α
                                             NO 2006-3057
     NO 2006003057
                          Δ
                                20060630
                                                                    20060630
PRIORITY APPLN. INFO.:
                                             EP 2003-104484
                                                                 Α
                                                                    20031202
                                             WO 2004-EP13683
                                                                 W
                                                                    20041201
OTHER SOURCE(S):
                         MARPAT 143:59817
     Title compds. A(YONO2)s [s = 1, 2; A = R1CH(OZ)CH2NZ1R2; R1 =
     1-naphthyloxymethyl, 4-(Me2CHOCH2CH2OCH2)C6H4OCH2, indol-4-yloxymethyl,
     carbazol-4-yloxymethyl, 4-MeSO2NHC6H4, etc.; R2 = CHMe2, CMe3,
     2-MeOC6H4OCH2CH2, etc.; Z = H, CO, CO2, etc.; Z1 = H, CO; Y = CO
     (substituted) alkylene, cycloalkylene, etc.], were prepared Thus,
     1-(9H-carbazol-4-yloxy)-3-[[2-(2-methoxyphenoxy)ethyl][(6-
     nitrooxyhexanoyl)amino]]-2-propanol (preparation from carvedilol and
     6-bromohexanoic acid described) increased cGMP levels in PC12 cells with
     EC50 = 0.6 \mu M.
     853906-63-5P 853906-72-6P
TΤ
     RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
        (claimed compound; preparation of nitrooxy derivs. of carvedilol and other
        β-blockers as antihypertensive drugs)
RN
     853906-63-5 CAPLUS
CN
     Butanoic acid, 4-(nitrooxy)-, 4-[3-[[3-(9H-carbazol-4-yloxy)-2-
     hydroxypropyl] [2-(2-methoxyphenoxy)ethyl]amino]-3-oxo-1-propenyl]-2-
     methoxyphenyl ester (9CI) (CA INDEX NAME)
```

PAGE 1-A

$$\begin{array}{c|c}
MeO
\end{array}$$

$$\begin{array}{c|c}
O_2N-O-(CH_2)_3-C-O\\
0
\end{array}$$

RN 853906-72-6 CAPLUS

CN Butanoic acid, 4-(nitrooxy)-, 4-[3-[2-(9H-carbazol-4-yloxy)-1-[[[3-[3-methoxy-4-[4-(nitrooxy)-1-oxobutoxy]phenyl]-1-oxo-2-propenyl][2-(2-methoxyphenoxy)ethyl]amino]methyl]ethoxy]-3-oxo-1-propenyl]-2-methoxyphenyl ester (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 2-A

CAPLUS COPYRIGHT 2007 ACS on STN L9 ANSWER 12 OF 98

1980:111045 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 92:111045

Quinazoline derivatives with antihypertensive TITLE:

action

Mizogami, Susumu; Hiranuma, Hidetoshi; Sekiya, Tetsuo; INVENTOR(S):

Hanazuka, Mitsuo

PATENT ASSIGNEE(S): Mitsubishi Yuka Yakuhin Co., Ltd., Japan

Ger. Offen., 66 pp. SOURCE:

CODEN: GWXXBX

DOCUMENT TYPE: Patent German LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|------|----------|------------------|----------|
| | | | | |
| DE 2848263 | A1 | 19790510 | DE 1978-2848263 | 19781107 |
| JP 54066691 | Α | 19790529 | JP 1977-133105 | 19771108 |
| US 4189484 | Α | 19800219 | US 1978-956326 | 19781031 |
| GB 2008106 | Α | 19790531 | GB 1978-43202 | 19781103 |
| ES 475173 | A1 | 19790416 | ES 1978-475173 | 19781107 |
| BE 871821 | A1 | 19790507 | BE 1978-191577 | 19781107 |
| DK 7804955 | A | 19790509 | DK 1978-4955 | 19781107 |
| SE 7811492 | A | 19790509 | SE 1978-11492 | 19781107 |
| NL 7811050 | A | 19790510 | NL 1978-11050 | 19781107 |
| FR 2407928 | A1 | 19790601 | FR 1978-31595 | 19781108 |
| PRIORITY APPLN. INFO.: | | | JP 1977-133105 A | 19771108 |
| | | | JP 1978-20891 A | 19780227 |

OTHER SOURCE(S):

MARPAT 92:111045

GΙ

The quinazoline derivs. I [R = (substituted) aryl, thienyl, furyl, or pyridyl; R1 = H, alkyl; n = 2, 3] were prepared for use as AB antihypertensives (test data tabulated). Thus, 2-chloro-4-amino-6,7-dimethoxyquinazoline reacted with 1-(4-methylcinnamoyl)piperazine to give I (R = 4-MeC6H4, R1 = H, n = 2). IT 33130-03-9 RL: RCT (Reactant); RACT (Reactant or reagent)

Ι

(reaction of, with piperazine)

RN 33130-03-9 CAPLUS

2-Propenoic acid, 3-(2,3,4-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

MeO
$$CH = CH - CO_2H$$

L9 ANSWER 13 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1996:666870 CAPLUS

DOCUMENT NUMBER: 125:301001

TITLE: Preparation of 3-(2'-sulfamoylbiphenyl-4-yl)methyl-2-

imino-1,3,4-thiazolidine derivatives as

antihypertensives

INVENTOR(S): Sakae, Shinya; Yokomoto, Masaharu; Inoe, Satoshi;

Nishimura, Koji; Hirata, Akikage; Iguma, Kenichi;

Tamura, Koichi

PATENT ASSIGNEE(S): Wakunaga Seiyaku Kk, Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 31 pp.

Patent

CODEN: JKXXAF

DOCUMENT TYPE:

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE | | |
|------------------------|-------|----------|-----------------|---|----------|--|--|
| JP 08208632 | А | 19960813 | JP 1995-280093 | | 19951027 | | |
| PRIORITY APPLN. INFO.: | | 13300013 | JP 1995-280093 | Α | 19951027 | | |
| | | | JP 1994-264755 | | 19941028 | | |

OTHER SOURCE(S): MARPAT 125:301001

GI

$$R^{3}$$
 $R^{1}N$
 N
 R^{5}
 $SO_{2}NHR^{4}$
 $Q=$
 $CO_{2}H$

AB The title compds. [I; R1 = H, COR2; wherein R2 = (un)substituted lower alkyl, cycloalkyl, or cycloalkenyl, (un)substituted aryl-lower alkyl or aryl-lower alkenyl, Ph, or aromatic heterocyclyl, lower alkoxy or aralkyloxy; R3 = halo, lower alkyl or cycloalkyl, (un) substituted Ph, lower alkyl alkoxy; R4 = H, lower alkyl, acyl; R5, R6 = H, halo, lower alkyl], which show potent angiotensin II-antagonizing, smooth muscle-relaxing, and antihypertensive activity, are prepared Thus, 533 mg 5-ethyl-2-trifluoroacetamido-1,3,4-thiadiazole and 1.00 g 4-bromomethyl-2'-(N-tert-butylsulfamoylbiphenyl-4-yl)biphenyl were added to DMF and stirred at room temperature for 4 h to give 606 mg I (R1 = CF3CO, R3 = Et, R5 = R6 = H, R4 = tert-butyl). I (R1 = Q, R3 = Et, R4 = CO2Et, R5 = R6 = H) and I (R1 = 2-C1C6H4CO, R3 = Et, R4 = COC6H4CO2Me-2, R5 = R6 = H) in vitro showed IC50 of 3.0 and 5.3 nM, resp., for inhibiting angiotensin II and in vivo inhibited angiotensin II-induced hypertension of rats by 53.4 and 62.3%, resp., at 0.1 mg/kg i.v.

IT 183000-48-8P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of [(sulfamoylbiphenylyl)methyl]iminothiazolidine derivs. as antihypertensives, angiotensin II antagonists, and smooth

muscle relaxants)

RN 183000-48-8 CAPLUS

CN Carbamic acid, [[4'-[[2-[[3-[3-(acetyloxy)-4-methoxyphenyl]-1-oxo-2-propenyl]imino]-5-ethyl-1,3,4-thiadiazol-3(2H)-yl]methyl][1,1'-biphenyl]-2-yl]sulfonyl]-, ethyl ester, (?,E)- (9CI) (CA INDEX NAME)

Double bond geometry as described by E or Z.

L9 ANSWER 14 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2006:362641 CAPLUS

DOCUMENT NUMBER:

144:350688

TITLE:

Losartan derivatives with antioxidant properties, and

their preparation and use as antihypertensives

with tissue damage prevention activities

INVENTOR(S):

Alajarin Ferrandez, Ramon; Alvarez-Builla Gomez, Julio; Diez Marques, Maria Luisa; Garcia Navazo, Gonzalo; Rodriguez Puyol, Diego; Rodriguez Puyol,

Manuel

1

PATENT ASSIGNEE(S):

Universidad de Alcala, Spain

SOURCE:

GΙ

Span., 19 pp. CODEN: SPXXAD

DOCUMENT TYPE:

Patent

LANGUAGE:

Spanish

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--------|---------------|--------------------------|----------|
| | | | | |
| ES 2242543 | A1 | 20051101 | ES 2004-1050 | 20040430 |
| PRIORITY APPLN. INFO.: | | | ES 2004-1050 | 20040430 |
| OTHER SOURCE(S): | CASREA | ACT 144:35068 | 88 · MARPAT 144 · 350688 | |

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Losartan derivs. I and a process for their preparation are disclosed [in which: X = H, Cl; A = residue of 8 specific phenolic carboxylic acid antioxidants, e.g., 3,4-dihydroxybenzoyl]. The preparation process involves Mitsunobu reaction of tritylated losartan derivative II with corresponding, optionally protected antioxidant acids, followed by appropriate deprotection of the obtained intermediate. Depending upon the deprotective conditions, the chlorine atom of II may or may not remain. I are prepared as pharmaceuticals with simultaneous angiotensin II

receptor-blocking and antioxidant properties, and are beneficial for preventing tissue damage in patients with cardiovascular risks. Thus, Mitsunobu reaction of II with 3-[3,4-bis(benzyloxy)phenyl]propanioic acid in the presence of PPh3 and di-Me azodicarboxylate in Et2O gave 63% intermediate III. Hydrogenolytic deprotection of III with 1 atm H2 over 30% Pd/C, with concomitant dechlorination, gave 56% invention compound IV, designated GGN 841. In tests for displacement of labeled angiotensin II from its receptor, and for inhibition of angiotensin II-induced contraction of human mesangial cells in vitro, IV was as active or slightly more active than losartan itself. In addition, the antioxidant activity of IV, determined by inhibition of the oxidation of ABTS in vitro, was 8-fold greater than that of losartan.

IT 331-39-5DP, 3,4-Dihydroxycinnamic acid, losartan derivative esters RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

> (drug candidate; losartan derivs. with antioxidant properties, and their preparation and use as antihypertensives with tissue damage prevention activities)

RN331-39-5 CAPLUS

2-Propenoic acid, 3-(3,4-dihydroxyphenyl)- (9CI) (CA INDEX NAME) CN

ANSWER 15 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1991:583950 CAPLUS

DOCUMENT NUMBER:

115:183950

TITLE:

Preparation of amino acid conjugates as

renal-selective prodrugs for the treatment of

hypertension

INVENTOR (S):

Reitz, David B.; Koepke, John P.; Blaine, Edward H.; Schuh, Joseph R.; Manning, Robert E.; Smits, Glenn J.

PATENT ASSIGNEE(S):

SOURCE:

G.D. Searle and Co., USA PCT Int. Appl., 459 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PA | PATENT NO. | | | | KIN | D DATE | APPLICATION NO. | | DATE |
|---------|------------|-------|-------|-----|-----|------------|------------------------|----|----------|
| WO | 9101 | 724 | | | A1 | 1991022 | NO 1990-US4168 | | 19900725 |
| | W: | CA, | JP, | KR, | US | | · | | |
| | RW: | AT, | BE, | CH, | DE, | DK, ES, FF | R, GB, IT, LU, NL, SE | | |
| EP | 48443 | 37 | | | A1 | 1992051 | L3 EP 1990-912307 | | 19900725 |
| | R: | AT, | BE, | CH, | DE, | DK, ES, FF | R, GB, IT, LI, LU, NL, | SE | |
| JP | 04506 | 5967 | | | T | 1992120 | JP 1990-511397 | | 19900725 |
| WO | 92016 | 567 | | | A1 | 1992020 | 06 WO 1991-US611 | | 19910128 |
| | W: | CA, | JΡ, | KR, | US | | | | |
| | RW: | AT, | BE, | CH, | DE, | DK, ES, FF | R, GB, GR, IT, LU, NL, | SE | |
| US | 20032 | 22052 | 21 | | A1 | 2003112 | 27 US 2002-151211 | | 20020520 |
| US | 2004 | 10152 | 23 | | A1 | 2004052 | 7 US 2003-689919 | | 20031020 |
| PRIORIT | Y APPI | LN. | INFO. | . : | | | US 1989-386527 | A2 | 19890727 |
| | | | | | | | WO 1990-US4168 | W | 19900725 |
| | | | | | | | US 1994-280170 | B1 | 19940725 |

US 1996-639493 B1 19960429 US 1999-444888 B1 19991122 US 2000-678015 A1 20001002 US 2002-151211 B1 20020520

OTHER SOURCE(S):

MARPAT 115:183950

GΙ

Bu
$$H$$
 CONNHCOCH₂CH₂CH (CO₂H) NHAC I

Title compds., conjugates comprising a 1st residue and a 2nd residue connected by a cleavable bond, wherein the 1st residue is an inhibitor of the biosynthesis of an adrenergic neurotransmitter and the 2nd residue is cleaved by an enzyme located predominantly in the kidney, are prepared 5-[(5-Butyl-2-pyridinyl)carbonyl]-L-glutamic acid hydrazide (preparation given) in MeCN/H2O was treated with 2 equiv of 1M K2CO3 followed by Ac2O and K2CO3 to give the L-glutamic hydrazide I. In spontaneously hypertensive rats, I at 8 mg/h lowered blood pressure from 146 to 122 mm Hg on day 1 and to 115 mm Hg on day 5. Addnl. compds. were prepared and tested. A large number of compds. are claimed.

TT 331-39-5DP, kidney enzyme-cleavable conjugate

IT 331-39-5DP, kidney enzyme-cleavable conjugate
RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of, as prodrug antihypertensive)

RN 331-39-5 CAPLUS

CN 2-Propenoic acid, 3-(3,4-dihydroxyphenyl)- (9CI) (CA INDEX NAME)

L9 ANSWER 16 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1994:671689 CAPLUS

DOCUMENT NUMBER:

121:271689

TITLE:

Caracasanamide - a novel antihypertensive

agent

AUTHOR (S):

Lee, An-Rong; Lin, Connie K.; Huang, Wen-Hsin; Chen,

Hsiu-Ho

CORPORATE SOURCE:

School Pharmacy, National Defense Medical Center,

Taipei, Taiwan

SOURCE:

Yixue Yanjiu (1994), 14(6), 357-70

CODEN: YIXYE3; ISSN: 1011-4564

DOCUMENT TYPE:

Journal

LANGUAGE:

Chinese

AB Caracasanamide is one of the hypotensive agents isolated from Verbesina caracasana. It is the mixture of E/Z forms of 1-[3,4-(dimethoxycinnamoyl)amino]-4-[(3-methyl-2-butenyl)guanidino]butane. At nontoxic dose, Z-caracasanamide possesses significant, in vivo, activity of antihypertension with a lasting duration. Z-Caracasanamide was found to decrease blood pressure and to increase cardiac inotropism, respiratory frequency, and tidal volume, and to induce a very slight and not significant tachycardia. The pharmacol. profiles indicate that the cardiovascular actions of this compound might result from: (a) a decrease in sympathetic outflow through central neurogenic mechanisms; (b) arterial vasodilation;

and, (c) an interaction with the cardiac β -1 adrenoreceptors. Z-Caracasanamide is as effective as reserpine in lowering blood pressure and demonstrates a longer duration than guanethidine, papaverine, histamine, and thus is a candidate in clin. application. This paper deals with the origin, structure, synthesis, pharmacol. actions and advantages of actions of caracasanamide.

IT 128009-16-5 146269-39-8, Caracasanamide

146269-40-1, z-Caracasanamide

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(antihypertensive activity and mechanism of action of)

RN 128009-16-5 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[[imino[(3-methyl-2-butenyl)amino]methyl]amino]butyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & \text{NH} \\ & \text{CH} \end{array} \begin{array}{c} \text{CH} = \text{CH} - \text{C-NH} - \text{(CH}_2)_4 - \text{NH} - \text{C-NH} - \text{CH}_2 - \text{CH} = \text{CMe}_2 \\ \\ \text{MeO} \end{array}$$

RN 146269-39-8 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[[imino[(3-methyl-2-butenyl)amino]methyl]amino]butyl]-, (E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 146269-40-1 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[[imino[(3-methyl-2-butenyl)amino]methyl]amino]butyl]-, (2Z)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

$$\begin{array}{c|c} & & & \\ & & \\ \hline & & \\ & & \\ \hline & & \\ & &$$

L9 ANSWER 17 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 20

2000:900158 CAPLUS

DOCUMENT NUMBER:

135:55773

TITLE:

Reduction in left ventricular messenger RNA for transforming growth factor $\beta 1$ attenuates left ventricular fibrosis and improves survival without lowering blood pressure in the hypertensive

TGR(mRen2)27 rat

AUTHOR (S):

SOURCE:

Pinto, Yigal M.; Pinto-Sietsma, Sara-Joan; Philipp, Tobias; Engler, Sonja; Kossmehl, Peter; Hocher, Berthold; Marquardt, Heike; Sethmann, Svenja; Lauster, Roland; Merker, Hans-Joachim; Paul, Martin

CORPORATE SOURCE:

Department of Clinical Pharmacology and Toxicology Benjamin Franklin Medical Center, Freie Universitat

Berlin, Berlin, 14195, Germany

Hypertension (2000), 36(5), 747-754

CODEN: HPRTDN; ISSN: 0194-911X Lippincott Williams & Wilkins

PUBLISHER: DOCUMENT TYPE:

Journal English

DOCUMENT TYPE: LANGUAGE:

Angiotensin II recruits transforming growth factor β 1 (TGF β 1) and is related to left ventricular fibrosis. However, it is unclear whether chronic in vivo reduction in left ventricular TGFB1 expression blunts fibrosis and improves outcome in angiotensin II-dependent hypertension. Four-week-old male hypertensive TGR(mRen2)27 (Ren2) rats received either normal food, low-dose losartan (0.5 mg/kg/d), or tranilast (a nonspecific TGF β inhibitor; 400 mg/kg/d) for 12 wk and were compared with Sprague-Dawley control rats. The effect of tranilast on survival was evaluated in 34 addnl. untreated homozygous Ren2 rats. Tranilast or low-dose losartan did not lower blood pressure. However, the increase in left ventricular weight (Ren2 vs. SD 3.1 vs. 2.1 mg/g) was significantly blunted by both tranilast (2.7) and losartan (2.7). Both drugs prevented the increase in left ventricular $TGF\beta 1$ mRNA and fibronectin mRNA and blunted the increase in hydroxyproline content and the increase in perivascular fibrosis. perivascular fibrosis score correlated significantly with the level of expression of $TGF\beta1$ (r = 0.62). In situ hybridization demonstrated increases in TGF\$1 mRNA, predominantly in perivascular and nonmyocyte areas. Both drugs did not prevent the decrease in systolic or diastolic dP/dt, but tranilast significantly improved the survival of untreated Ren2 In conclusion, TGFβ1 mRNA expression is increased predominantly in nonmyocyte regions in the hypertrophied left ventricle in this angiotensin II-dependent model of hypertension. This increase is probably due to high angiotensin II levels rather than to hypertension. This is the first study to suggest that chronic inhibition of TGFβ1 expression attenuates left ventricular hypertrophy and fibrosis, even without lowering blood pressure. ΙT 53902-12-8, Tranilast

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(TGF- β 1 mRNA reduction in left ventricle attenuates left ventricular fibrosis and improves survival without lowering blood pressure in hypertensive TGR(mRen2)27 rats)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

REFERENCE COUNT: 35 THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 18 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN L9

ACCESSION NUMBER: 2004:113676 CAPLUS

DOCUMENT NUMBER:

141:64675

TITLE:

Effects of sodium ferulate on plasma levels of

endothelin-1 and nitric oxide in patients with renal

hypertension and chronic renal insufficiency

Yang, Jinghua; Zhou, Qiaoling; Cheng, Xiaomiao; Deng,

Shengli; Wu, Cailing

CORPORATE SOURCE:

Department of Nephrology, Xiangya Hospital, Central South University, Changsha, 410008, Peop. Rep. China

SOURCE:

Hunan Yike Daxue Xuebao (2002), 27(5), 445-447

CODEN: HYXBET; ISSN: 1000-5625

PUBLISHER:

AUTHOR (S):

Hunan Yike Daxue

Journal Chinese

DOCUMENT TYPE: LANGUAGE:

The changes of plasma endothelin-1 (ET-1) and nitric oxide and the effect of Na ferulate on patients with renal hypertension and chronic renal insufficiency were studied. Group I: sixty patients with renal hypertension and chronic renal insufficiency were divided into two groups: A and B. The patients in Group A were treated with Na ferulate and the routine therapy while those in Group B were treated only with the routine therapy. The serum concns. of ET-1 and NO were measured. The level of plasma ET-1 was higher and the level of NO was lower in Group A and B than those in Group C (P < 0.01). In Group A, plasma ET-1, blood urea-N (BUN), creatinine, and urinary protein were decreased while plasma NO was increased significantly after the treatment (P <0.01). Compared with Group B, those changes in Group A were more significant (P <0.01). There was a pos. correlation between ET-1 and blood pressure (Bp). There was a neg. correlation between NO and Bp. The level of plasma ET-1 of the patients was remarkably higher than that of the normal subjects in the control group, while NO was remarkably lower; Na ferulate can regulate the balance of plasma ET-1 and NO in patients with renal hypertension and chronic renal insufficiency; and Na ferulate played an important role in protecting renal functions and delaying chronic renal failure.

IT24276-84-4, Sodium ferulate

RL: DMA (Drug mechanism of action); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(effects of sodium ferulate on plasma levels of endothelin-1 and nitric oxide in patients with renal hypertension and chronic renal insufficiency)

24276-84-4 CAPLUS RN

2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)-, monosodium salt (9CI) CN (CA INDEX NAME)

Na

ANSWER 19 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2004:514574 CAPLUS

DOCUMENT NUMBER:

141:46677

TITLE: Tranilast and hypertensive heart disease:

Further insights into mechanisms of an anti-inflammatory and anti-fibrotic drug

AUTHOR(S): Pfab, Thiemo; Hocher, Berthold

CORPORATE SOURCE: Center for Cardiovascular Research (CCR) and

Department of Nephrology, Berlin, Germany

SOURCE: Journal of Hypertension (2004), 22(5), 883-886

CODEN: JOHYD3; ISSN: 0263-6352 Lippincott Williams & Wilkins

PUBLISHER: Lippincott Williams & W DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

AB A review on pharmacol. characteristics of translast, anti-proliferative

and antifibrotic effects (apart from mast-cell stabilization), and

anti-inflammatory potency of tranilast, mol. mechanisms of

anti-inflammatory tranilast action, pathogenesis of hypertensive

heart disease, and preventive effect of tranilast on cardiac fibrosis. A

polemic with S. Kagitani et al. (ibid. 2004, 22, 1007) is added,

concerning anti-inflammatory aspects of translast action in the course of

cardiac fibrosis development. Tranilast is supposed to be a promising compound against fibrotic remodeling of the heart and possibly other organs

such as the kidney in diabetic nephropathy.

IT 53902-12-8, Tranilast

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU

(Therapeutic use); BIOL (Biological study); USES (Uses)

(mol. mechanisms of antiinflammatory and antifibrotic effects of

tranilast in hypertensive heart disease and heart fibrosis)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

REFERENCE COUNT: 49 THERE ARE 49 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 20 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2001:581843 CAPLUS

DOCUMENT NUMBER: 135:180762

TITLE: Preparation of n

Preparation of nitrogen-containing compounds having

kinase inhibitory activity and drugs-containing the

same

INVENTOR(S): Takami, Atsuya; Iijima, Hiroshi; Iwakubo, Masayuki;

Okada, Yuji

PATENT ASSIGNEE(S): Kirin Beer Kabushiki Kaisha, Japan

SOURCE: PCT Int. Appl., 372 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

WO 2001056988 A1 20010809 WO 2001-JP721 20010201

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,

HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG AU 2001-30564 20010814 20010201 AU 2001030564 Α5 EP 2001-902730 20021113 EP 1256574 Α1 20010201 AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR US 2004102437 Α1 20040527 US 2003-181943 20030519 JP 2000-24292 PRIORITY APPLN. INFO.: А 20000201 WO 2001-JP721 W 20010201

OTHER SOURCE(S):

MARPAT 135:180762

GΙ

AB Title compds. [HetXZ; Het = monocyclic heterocycle or dicycle heterocycle having at least one nitrogen; X = NHCONHQ, NHCOQ1; Q, Q1 independently = bond, alkylene, alkenylene; Z = H halo, monocyclohydrocarbon, dicyclohydrocarbon, tricyclohydrocarbon, heterocycle], pharmaceutically acceptable salts thereof and solvates of the same are prepared as Rho kinase inhibitors. Thus, the title compound I was prepared and biol. tested for blood presure lowering effect in spontaneous hypertensive rats and diminished urine protein excretion effect in rabbits having GBM-antibody-mediated kidney disease.

IT 353539-62-5P 353539-74-9P 353539-79-4P 353539-84-1P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(preparation of nitrogen-containing compds. having kinase inhibitory
activity)

RN 353539-62-5 CAPLUS

CN 2-Propenamide, 3-(3,4-dihydroxyphenyl)-N-1H-indazol-5-yl-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 353539-74-9 CAPLUS

CN 2-Propenamide, 3-(4-hydroxy-3-methoxyphenyl)-N-1H-indazol-5-yl-, (2E)-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

HO
$$E$$
 N N N

RN 353539-79-4 CAPLUS

CN 2-Propenamide, 3-(3-hydroxy-4-methoxyphenyl)-N-1H-indazol-5-yl-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 353539-84-1 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-1H-indazol-5-yl-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

$$\begin{array}{c} \text{OMe} \\ \\ \text{E} \\ \\ \text{O} \\ \end{array}$$

IT 501-16-6 537-98-4 14737-89-4

25522-33-2

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of nitrogen-containing compds. having kinase inhibitory activity)

RN 501-16-6 CAPLUS

CN 2-Propenoic acid, 3-(3,4-dihydroxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 537-98-4 CAPLUS

CN 2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 14737-89-4 CAPLUS

CN 2-Propenoic acid, 3-(3,4-dimethoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 25522-33-2 CAPLUS

CN 2-Propenoic acid, 3-(3-hydroxy-4-methoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

REFERENCE COUNT:

236 THERE ARE 236 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 21 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1990:429237 CAPLUS

DOCUMENT NUMBER:

113:29237

TITLE:

Guanidine derivatives having hypotensive activity, composition containing them, and process for obtaining

them

INVENTOR(S):

Delle Monache, Giuliano; Delle Monache, Franco; Botta,

Bruno; Bonnevaux Castillo, Stella; Espinal, Romulo; De

Luca, Carlo; Carmignani, Marco

PATENT ASSIGNEE(S): Consiglio Nazionale delle Ricerche, Italy

SOURCE: Eur. Pat. Appl., 26 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--------|--------------|----------------------|----------|
| | | | | |
| EP 330629 | A2 | 19890830 | EP 1989-830067 | 19890217 |
| EP 330629 | A3 | 19901031 | | |
| R: AT, BE, CH, | DE, ES | , FR, GB, GR | , IT, LI, LU, NL, SE | |
| JP 02003661 | Α | 19900109 | JP 1989-45003 | 19890223 |
| US 5059624 | Α | 19911022 | US 1989-315107 | 19890224 |
| PRIORITY APPLN. INFO.: | | | IT 1988-47665 A | 19880224 |
| OTHER SOURCE(S): | MARPAT | 113:29237 | • | • |
| GT | | | | |

AB Guanidine derivs. R1NHCH2NHC:NHNHR2, and R2NHC:NHNH(CH2)nNHR3NH(CH2)nNHC:N HNHR2 [R1 = H, (substituted)cinnamoyl; R2 = H, alkyl, alkenyl with the proviso both R1 and R2 ≠ H; R3 = (substituted)truxinoyl, (substituted)truxilloyl; n = 1-8], useful as hypotensives, may be synthesized or isolated from Verbesina caracasana. Thus, V. caracasana was extracted with MeOH, the residue was extracted with EtOAc/H2O, and the H2O-soluble portion was lyophilized and then resuspended in MeOH. Chromatog. of the solution on silica using CHCl3 eluant yielded I 3-6 g. I, administered i.v. to rats at 50-6400 μg/kg, reduced arterial pressure and increased heart rate and respiration rate. I LD50 i.p. in mice was 57 mg/kg. Synthesis of I is also described.

IT 128009-20-1

RL: PROC (Process)

(isolation of, from extract of Verbesina caracasana)

RN 128009-20-1 CAPLUS

CN 2-Propenamide, N-[4-[(aminocarbonyl)amino]butyl]-3-(3,4-dimethoxyphenyl)(9CI) (CA INDEX NAME)

IT 128009-16-5 128009-18-7

RL: PROC (Process)

(isolation of, from Verbesina caracasana, as antihypertensive

128009-16-5 CAPLUS RN

2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[[imino[(3-methyl-2-CNbutenyl)amino]methyl]amino]butyl]- (9CI) (CA INDEX NAME)

RN128009-18-7 CAPLUS

CN2-Propenamide, N-[4-[(aminoiminomethyl)amino]butyl]-3-(3,4dimethoxyphenyl) - (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} \text{O} & \text{NH} \\ \parallel & \parallel \\ \text{CH-C-NH-(CH_2)_4-NH-C-NH_2} \\ \\ \text{MeO} & \text{OMe} \end{array}$$

128009-24-5P IT

RL: RCT (Reactant); PREP (Preparation); RACT (Reactant or reagent) (preparation and decarboxylation of, in antihypertensive preparation)

RN128009-24-5 CAPLUS

CN Carbamic acid, [[4-[[3-(3,4-dimethoxyphenyl)-1-oxo-2propenyl]amino]butyl]carbonimidoyl]bis-, bis(1,1-dimethylethyl) ester (9CI) (CA INDEX NAME)

ANSWER 22 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1999:659729 CAPLUS

DOCUMENT NUMBER: 131:295291

TITLE: Effect of tranilast on the retinal vessels in the

hypertensive rat

AUTHOR (S): Honda, Yukie; Aoike, Chiaki

CORPORATE SOURCE: Second Dep. Ophthalmol., Toho Univ. Sch. Med., 2-17-6

Ohashi, Meguro-ku, Tokyo, 153-0044, Japan

SOURCE: Atarashii Ganka (1999), 16(9), 1291-1294

CODEN: ATGAEX; ISSN: 0910-1810 PUBLISHER:

Medikaru Ai Shuppan

DOCUMENT TYPE: Journal LANGUAGE: Japanese

We investigated the effect of tranilast on the retinal vessels in spontaneously hypertensive rats (SHR). Salt loading stroke-prone SHR (SHR-sp) were assigned to either the treated group (dosed with tranilast) or the untreated group. After treatment, computer imaging anal. showed retinal vessel thickness to be significantly inhibited in the treatment group after 8 wk of treatment (p = 0.008). This result suggests that tranilast may have an inhibitory effect on early stage hypertensive retinopathy.

53902-12-8, Tranilast TΤ

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES

(effect of translast on retinal vessels in hypertensive rat)

RN53902-12-8 CAPLUS

Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) CN (CA INDEX NAME)

ANSWER 23 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN L9

ACCESSION NUMBER: 1973:72199 CAPLUS

DOCUMENT NUMBER:

78:72199

TITLE:

Pharmacologically active acyl derivatives of

1-aminopiperazines

INVENTOR(S):

Fauran, C.; Turin, M.; Raynaud, G.; Dorme, N. Delalande S. A.

PATENT ASSIGNEE(S): SOURCE:

Fr. Demande, 14 pp.

CODEN: FRXXBL

DOCUMENT TYPE:

Patent

LANGUAGE:

French

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| · | PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|------|--------------------------|----------|----------------------|----------------------|-----|--------------|
| | FR 2115024 FR 2115024 | A5 B1 | 19720707 19740322 | FR 1970-42063 | _ | 19701124 |
| PRIO | RITY APPLN. INFO.: | | | FR 1970-42063 | Α | 19701124 |
| GI | For diagram(s), see | | | | | |
| AB | | | | azines I [R = NHCHMe | | pyrrolidino; |
| | | | | CH, 3,4,5-(MeO)3C6H2 | | |
| | | | | H:CH, 3,4-(MeO)2C6H3 | | |
| | | | | re prepared by acyla | | |
| | | | | es. I exhibited hyp | | |
| | | | analgesic, | antiinflammatory, ch | ole | retic, and |
| | spasmolytic activit | • | | | | |
| IT | 39855-73-7P 39855-7 | | | | | |
| | 39855-82-8P 39855-8 | 7-3P 39 | 855-97 - 5P | | | |
| | 39855-98-6P | | | | | |
| | RL: SPN (Synthetic | prepara | tion); PREP | (Preparation) | | |

(preparation of)

RN 39855-73-7 CAPLUS

CN 2-Propenamide, N-[4-[2-oxo-2-(1-pyrrolidinyl)ethyl]-1-piperazinyl]-3-(3,4,5-trimethoxyphenyl) - (9CI) (CA INDEX NAME)

RN 39855-74-8 CAPLUS

CN 2-Propenamide, N-[4-[2-oxo-2-(1-pyrrolidinyl)ethyl]-1-piperazinyl]-3-(3,4,5-trimethoxyphenyl)-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 39855-73-7 CMF C22 H32 N4 O5

CM 2

CRN 110-16-7 CMF C4 H4 O4

Double bond geometry as shown.

RN 39855-81-7 CAPLUS

CN 1-Piperazineacetamide, N-(1-methylethyl)-4-[[1-oxo-3-(3,4,5-trimethoxyphenyl)-2-propenyl]amino]- (9CI) (CA INDEX NAME)

RN 39855-82-8 CAPLUS

CN 1-Piperazineacetamide, N-(1-methylethyl)-4-[[1-oxo-3-(3,4,5-trimethoxyphenyl)-2-propenyl]amino]-, monohydrochloride (9CI) (CA INDEX NAME)

HCl

RN 39855-87-3 CAPLUS

CN 1-Piperazineacetamide, 4-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]-N-(1-methylethyl)- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} \text{O} & \text{OMe} \\ \text{i-PrNH-C-CH}_2 & \text{OMe} \\ \\ \text{N-NH-C-CH} & \text{CH-CH} \end{array}$$

RN 39855-97-5 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[2-oxo-2-(1-pyrrolidinyl)ethyl]-1-piperazinyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & \\ & & \\ \text{MeO} & & \\ &$$

RN 39855-98-6 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[2-oxo-2-(1-pyrrolidinyl)ethyl]-1-piperazinyl]-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CM 1

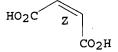
CRN 39855-97-5 CMF C21 H30 N4 O4

MeO CH CH CH CH NH N
$$\sim$$
 CH₂ C \sim N \sim CH₂ C \sim N

CM 2

CRN 110-16-7 CMF C4 H4 O4

Double bond geometry as shown.



L9 ANSWER 24 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1977:106677 CAPLUS

DOCUMENT NUMBER: 86:106677

TITLE: Piperazine- and homopiperazinealkanol esters

INVENTOR(S): Kato, Hideo; Nishikawa, Tomoyasu; Mouri, Takaaki

PATENT ASSIGNEE(S): Hokuriku Pharmaceutical Co., Ltd., Japan

SOURCE: Austrian, 11 pp. CODEN: AUXXAK

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|------|----------|-----------------|------------|
| | | | · | |
| AT 333287 | В | 19761110 | AT 1974-135 | 19740109 |
| AT 7400135 | Α | 19760315 | | |
| JP 50062983 | A | 19750529 | JP 1973-112289 | 19731008 |
| JP 52024031 | В | 19770628 | | |
| JP 50062988 | A | 19750529 | JP 1973-112290 | 19731008 |
| JP 52046235 | В | 19771122 | | |
| JP 50062984 . | Α | 19750529 | JP 1973-112291 | 19731008 |
| JP 52024032 | В | 19770628 | | |
| PRIORITY APPLN. INFO.: | | | JP 1973-112289 | A 19731015 |
| | | | JP 1973-112290 | A 19731015 |
| | | | JP 1973-112291 | A 19731015 |

GΙ

$$N = N (CH_2)_{m}O_2CR$$

AB Hexahydro-1H-1,4-diazepine- and piperazine-1-alkanol esters [I; R = e.g. 2,3,4-(MeO) 3C6H2, 3,4,5-(MeO3) 3C6H2, 3,4-(MeO) 2C6H3, 2-ClC6H4, Ph, 2-pyridinyl, PhCH:CH; n = 1, 2; m = 2, 3], useful as antihypertensives (no data), are prepared by esterification of the alkanols with the appropriate acids. Thus, refluxing of 1-piperazineethanol and 2,3,4-(MeO) 3C6H2CO2H in presence of 4-MeC6H4SO3H 15 h in C6H6 and treatment with maleic acid gives 46% I dimaleate monohydrate [R = 2,3,4-(MeO) 3C6H2, n = 1, m = 2].

IT 90-50-6 33130-03-9

RL: RCT (Reactant); RACT (Reactant or reagent)

(esterification by, of piperazine- and homopiperazinealkanols)

RN 90-50-6 CAPLUS

CN 2-Propenoic acid, 3-(3,4,5-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

RN 33130-03-9 CAPLUS

CN 2-Propenoic acid, 3-(2,3,4-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

MeO CH CH
$$\sim$$
 CH \sim C

L9 ANSWER 25 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1980:94421 CAPLUS

DOCUMENT NUMBER: 92:94421

TITLE: Alkylenediamine derivatives

INVENTOR(S): Philippe, Michel; Manoury, Jacques

PATENT ASSIGNEE(S): Synthelabo S. A., Fr.

SOURCE: Fr. Demande, 12 pp. Addn. to Fr. Demande 2,362,630.

CODEN: FRXXBL

DOCUMENT TYPE: Patent LANGUAGE: French

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--------|----------|-----------------|----------|
| | | | | |
| FR 2389613 | A2 | 19781201 | FR 1977-13659 | 19770505 |
| FR 2389613 | B2 | 19801205 | | |
| PRIORITY APPLN. INFO.: | | | FR 1977-13659 A | 19770505 |
| OTHER SOURCE(S): | MARPAT | 92:94421 | | |
| GI | | | • | |

$$\underset{\text{MeO}}{\text{MeO}} \xrightarrow{N} \underset{\text{NR-C}_{n}\text{H}_{2n}\text{-NHR}^{1}} \\ \underset{\text{NH}_{2}}{\text{NR-C}_{n}\text{H}_{2n}\text{-NHR}^{1}}$$

The diamines I [R,R1(same or different) = H, C1-5 alkyl; R2 = (un)substituted phenyl, C1-4 alkoxy, alkyl, pyridyl, furyl, etc.; n = 2-4; Y = CO, SO2], having relatively long-lived antihypertensive activity, were prepared by condensing II (X = halo) with RNHCnH2nNR1YR2 or III with XYR2. Thus, stirring I (R = R1 = Me, n = 3) with

III

 $3,4,5-(MeO)\ 3C6H2CH:CHCOCl$ in CHCl3 at room temperature 30 min gave 47% I [R =

R1

= Me, n = 3, R2 = 3,4,5-(MeO)3C6H2CH:CH, <math>Y = CO], isolated as HCl salt.

IT 72766-58-6P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(preparation and antihypertensive activity of)

RN 72766-58-6 CAPLUS

CN 2-Propenamide, N-[3-[(4-amino-6,7-dimethoxy-2-quinazoliny1)methylamino]propy1]-N-methyl-3-(3,4,5-trimethoxypheny1)-, monohydrochloride (9CI) (CA INDEX NAME)

MeO N N N (CH₂)
$$_3$$
 N C CH CH OMe OMe OMe

⊕ HCl

L9 ANSWER 26 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1965:11279 CAPLUS

DOCUMENT NUMBER:

62:11279

ORIGINAL REFERENCE NO.:

62:2093g-h,2094a

TITLE:

SOURCE:

Urinary excretion of phenolic acids and indole

derivatives in hypertonias

AUTHOR(S):

Borschel, W.; Hartmann, F.; Heimsoth, V.; Ruge, W.

CORPORATE SOURCE:

Med. Univ. Poliklin., Marburg/Lahn, Germany Klinische Wochenschrift (1964), 42(19), 927-35

CODEN: KLWOAZ; ISSN: 0023-2173

DOCUMENT TYPE:

Journal

LANGUAGE:

German

Using paper chromatog., .apprx.40 phenolic acids and .apprx.35 indole and other Ehrlichpos. substances were demonstrated in the urine of patients having various forms of hypertension, and of normals. Of these, 26 phenolic acids and .apprx.20 indole, etc. derivs. were identified as known metabolic products. Qual. changes in the excretion pattern of hypertensive, as compared to normal, urine occurred only in the case of m-hydroxybenzoic acid, which was present only in urines of hypertensive patients. Excretion of p-hydroxybenzoic, o-hydroxyhippuric, 3-(4-hydroxy-3-methoxyphenyl)hydracrylic acid, and vanilmandelic acid was increased irregularly in hypertensive urines. Vanillic acid excretion was increased in some patients, and diminished in others. The most distinct changes compared with normal urines appeared in a group of juvenile hypertensives with highly labile blood pressure values. There were no changes in the excretion of tryptophan-degradation products. 25 refs.

IT 1220-05-9, Glycine, N-(4-hydroxy-3-methoxycinnamoyl)-

(in urine chromatog. of, in hypertension)

RN 1220-05-9 CAPLUS

CN Glycine, N-[3-(4-hydroxy-3-methoxyphenyl)-1-oxo-2-propenyl]- (9CI) (CA INDEX NAME)

1135-24-6, Cinnamic acid, 4-hydroxy-3-methoxy-IT (in urine, in hypertension)

1135-24-6 CAPLUS RN

2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)- (9CI) (CA INDEX NAME) CN

$$CH = CH - CO_2H$$
OMe

L9 CAPLUS COPYRIGHT 2007 ACS on STN ANSWER 27 OF 98

ACCESSION NUMBER: 1983:498801 CAPLUS

DOCUMENT NUMBER:

99:98801

TITLE:

Preparation and study of 3-phenyl-3hydroxyaminopropionhydroxamic acids

AUTHOR (S):

Fountain, K. R.; Early, T.; Erwin, R.; Aijaz, S.;

Kehl, H.

CORPORATE SOURCE:

SOURCE:

Northeast Missouri State Univ., Kirksville, MO, USA Chem. Biol. Hydroxamic Acids, [Proc. Int. Symp.], 1st (1982), Meeting Date 1981, 51-62. Editor(s): Kehl,

Horst. Karger: Basel, Switz.

CODEN: 49RQAC

Ι

II

DOCUMENT TYPE:

LANGUAGE:

Conference English

GI

$$R^{2}$$
 CH (NHOH) $CH_{2}CONHOH$

$$R^1$$
—CH=CHCONR³OH

AB The hydroxamic acids I (R1 = H, NO2, -CH2O, OMe, or Me; R2 = H, Cl, Br, -CH2O, F, or OMe; R3 = H or C1), II (R1 = R2 = H or OMe; R3 = H or Me), and PhCH(NROH)CH2CONHOH (R = Me, Et, n-heptyl, 3-pyridyl, or hydrocinnamyl) were prepared and some tested for blood pressure-lowering activity in normotensive dogs. Structure-activity relations are discussed. Among the I derivs., for example, the m-Cl (R1 = Cl; R2 = R3 =[67248-11-7] and the m-F (R1 = F; R2 = R3 = H) [67248-13-9] derivs. were the most potent. The 4-nitro derivative (R1 = R3 = H; R2 = NO2) [86933-61-1] was the least active, and the remaining compds. were intermediate in potency.

IT 86933-56-4P

> RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation and antihypertensive activity of)

RN86933-56-4 CAPLUS

2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-hydroxy-N-methyl-, (E)- (9CI) CN(CA INDEX NAME)

Double bond geometry as shown.

Ь9 ANSWER 28 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1996:190216 CAPLUS

DOCUMENT NUMBER: 124:331677

TITLE: Hypotensive agents from Verbesina caracasana. 4.

Synthesis and preliminary pharmacological evaluation of analogs of caracasanamide, a hypotensive natural

product

Corelli, Federico; Dei, Donata; Delle Monache, AUTHOR (S):

Giuliano; Botta, Bruno; De Luca, Carlo; Carmignani,

Marco; Volpe, Anna Rita; Botta, Maurizio

Ι

CORPORATE SOURCE:

Dipartimento Farmaco Chimico Technologico, Universita

Siena, Siena, 53100, Italy

SOURCE: Bioorganic & Medicinal Chemistry Letters (1996), 6(6),

653-8

CODEN: BMCLE8; ISSN: 0960-894X

PUBLISHER: Elsevier DOCUMENT TYPE: Journal LANGUAGE: English

OTHER SOURCE(S): CASREACT 124:331677

GI

$$CH_2CH = CHNH (CH_2)_4NH$$
 $NHCH_2CH = CMe_2$
 HN
@ MeSO₃H

Some analogs of the hypotensive agent caracasanamide have been synthesized and tested in vivo for cardiovascular effects. The effects of the compds. on the heart and respiratory effects were also determined I emerged as the most interesting compound in the series. Structure-activity relation is also discussed.

IT 176640-44-1P 176640-45-2P 176640-46-3P

176640-47-4P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(synthesis and preliminary pharmacol. evaluation of analogs of caracasanamide as hypotensive agents in relation to structure)

RN176640-44-1 CAPLUS

2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[5-[[imino[(3-methyl-2-CN butenyl)amino]methyl]amino]pentyl]-, (E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

MeO
$$E$$
 CMe_2 MeO H H H H

RN 176640-45-2 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[5-[[imino[(3-methyl-2butenyl)amino]methyl]amino]pentyl]-, (2E)-, monomethanesulfonate (9CI) (CA INDEX NAME)

CM 1

CRN 176640-44-1 CMF C22 H34 N4 O3

Double bond geometry as shown.

MeO
$$E$$
 $(CH_2)_5$ NH NH NH MH MH

CM 2

CRN 75-75-2 CMF C H4 O3 S

RN 176640-46-3 CAPLUS

2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[6-[[imino[(3-methyl-2-CN butenyl)amino]methyl]amino]hexyl]-, (E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

MeO
$$E$$
 CMe_2 MeO H H H H

RN 176640-47-4 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[6-[[imino[(3-methyl-2-butenyl)amino]methyl]amino]hexyl]-, (2E)-, monomethanesulfonate (9CI) (CA INDEX NAME)

CM 1

CRN 176640-46-3 CMF C23 H36 N4 O3

Double bond geometry as shown.

MeO
$$E$$
 CMe_2 MeO H H H H H

CM 2

CRN 75-75-2 CMF C H4 O3 S

IT 146269-40-1, (Z)-Caracasanamide

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(synthesis and preliminary pharmacol. evaluation of analogs of caracasanamide as hypotensive agents in relation to structure)

RN 146269-40-1 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[[imino[(3-methyl-2-butenyl)amino]methyl]amino]butyl]-, (2Z)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

$$\begin{array}{c|c} & H & H & H \\ \hline Z & O & (CH_2)_4 & NH & N \\ \hline \end{array}$$

L9 ANSWER 29 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1986:186376 CAPLUS

DOCUMENT NUMBER: 104:186376

TITLE: Synthesis of N,N'-bis(ω -

aroylamidoalkyl)piperazine derivatives

AUTHOR(S): Sun, Qingfen; Zhu, Cuili

CORPORATE SOURCE: Shanghai 1st Med. Coll., Fac. Pharm., Shanghai, Peop.

Rep. China

Ι

SOURCE: Shanghai Diyi Yixueyuan Xuebao (1985), 12(3), 178-82

CODEN: SIIPD4; ISSN: 0253-3650

DOCUMENT TYPE: LANGUAGE:

: Journal Chinese

GΙ

RCOHN (CH₂)_nN (CH₂)_nNHCOR

The title compds. I [R = 3,4-(MeO)2C6H3, 3,4,5-(MeO)3C6H2, 3,4,5-(MeO)3C6H2CH:CH, 3,4-(MeO)2C6H3CH:CH, 4-MeOC6H4CH:CH, R1C6H4; R1 = 2-, 4-MeO, 4-H2N, 4-Br, 4-Cl, n = 2-3] were prepared by amidation of RCOCl with N,N'-bis(aminoalkyl)piperazines, obtained from alkylation, followed by catalytic hydrogenation of piperazine with CH2:CHCN or ClCH2CN. I (R = 4-MeOC6H4CH:CH) showed similar hypotensive activity as dilazep.

IT 101913-56-8P 101913-57-9P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation and hypotensive activity of)

RN 101913-56-8 CAPLUS

CN 2-Propenamide, N,N'-(1,4-piperazinediyldi-3,1-propanediyl)bis[3-(3,4,5-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

PAGE 2-A

RN 101913-57-9 CAPLUS

CN 2-Propenamide, N,N'-(1,4-piperazinediyldi-3,1-propanediyl)bis[3-(3,4-dimethoxyphenyl)- (9CI) (CA INDEX NAME)

PAGE 2-A

L9 ANSWER 30 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2007:13633 CAPLUS

DOCUMENT NUMBER:

146:121747

TITLE:

Prostaglandin derivatives

INVENTOR(S):

Benedini, Francesca; Chiroli, Valerio; Chong, Wesley

Kwan Mung; Krauss, Achim; Niesman, Michael Ross;

Ongini, Ennio

PATENT ASSIGNEE(S):

Pfizer Inc., USA; Nicox S.A.

SOURCE:

PCT Int. Appl., 110pp.

DOCUMENT TYPE:

CODEN: PIXXD2

LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PAT | PATENT NO. | | | | KIND DATE | | | APPLICATION NO. | | | | | | DATE | | | |
|----------------|------------|-----|------|-------------|-----------|-----|-----|-----------------|------|------|------|------|----------|------|------|------|-----|
| | | | | | | | | | | | | | | | | | |
| WO | 2007000641 | | | A2 20070104 | | | 1 | WO 2 | 006- | IB17 | 27 | | 20060619 | | | | |
| | W: | ΑE, | AG, | AL, | AM, | ΑT, | ΑU, | ΑZ, | BA, | BB, | BG, | BR, | BW, | BY, | ΒZ, | CA, | CH, |
| | | CN, | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EC, | EE, | EG, | ES, | FI, | GB, | GD, |
| | | GE, | GH, | GM, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | KE, | KG, | KM, | KN, | ΚP, | KR, |
| | | ΚZ, | LA, | LC, | LK, | LR, | LS, | LT, | LU, | LV, | LY, | MA, | MD, | MG, | MK, | MN, | MW, |
| | | MX, | ΜZ, | NA, | NG, | NI, | NO, | NZ, | OM, | PG, | PH, | PL, | PT, | RO, | RS, | RU, | SC, |
| | | SD, | SE, | SG, | SK, | SL, | SM, | SY, | ТJ, | TM, | TN, | TR, | TT, | TZ, | UA, | UG, | US, |
| | | UΖ, | VC, | VN, | ZA, | ZM, | ZW | | | | | | | | | | |
| | RW: | ΑT, | BE, | BG, | CH, | CY, | CZ, | DE, | DK, | EE, | ES, | FI, | FR, | GB, | GR, | HU, | ΙE, |
| | | IS, | IT, | LT, | LU, | LV, | MC, | NL, | PL, | PT, | RO, | SE, | SI, | SK, | TR, | BF, | ВJ, |
| | | CF, | CG, | CI, | CM, | GΑ, | GN, | GQ, | GW, | ML, | MR, | NE, | SN, | TD, | TG, | BW, | GH, |
| | | GM, | ΚE, | LS, | MW, | MZ, | NΑ, | SD, | SL, | SZ, | TZ, | ŪĠ, | ZM, | ZW, | AM, | ΑZ, | BY, |
| | | KG, | KZ, | MD, | RU, | ТJ, | TM | | | | | | | | | | |
| PRIORITY GI | APP | LN. | INFO | . : | | ÷ | | | Ţ | JS 2 | 005- | 6963 | 83P | , | P 20 | 0050 | 629 |

AB Nitrooxy derivs. of prostaglandin amides, such as I [R = NH-X-ONO2, NH-X = amide linking group in which X may be alkylene, arylene, alkenylene, ether, thioether or NH-X is an amino acid residue or a combination thereof; R1 = CH2Ph, OPh, OC6H4-3-CF3, OC6H4-3-Cl, (CH2)5Me; 13,14-bond = (E)-double or single; R15a = OH, R15b = H or R15aR15b = O], with improved pharmacol. activity and enhanced tolerability were prepared for therapeutic use in ophthalmic compns. for the treatment of glaucoma and ocular hypertension. Thus, prostaglandin amide II was prepared via an amidation reaction of the bis-O-(tert-butyltrimethylsilyl) protected derivative of latanoprost acid with the hydrobromide salt of Br(CH2)3NH2 using TEA, EDAC and DMAP in CH2Cl2, conversion of the resulting brominated amide to its nitrooxy derivative using AgNO3 in MeCN, and finally, desilylation of the resulting nitrooxy derivative using TBAF in THF. The prepared prostaglandin

II

amides were assayed in rabbits for their effect on hypertonic saline-induced transient intraocular pressure rise.

IT 537-98-4

HO

НО

RL: RCT (Reactant); RACT (Reactant or reagent)
(preparation of prostaglandin derivs. containing a nitrooxy moiety for

therapeutic use in the treatment of glaucoma and ocular hypertension)

RN 537-98-4 CAPLUS

CN 2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L9 ANSWER 31 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1974:520694 CAPLUS

DOCUMENT NUMBER: 81:120694

TITLE: 1-(Hydroxyalkyl)piperazine and -homopiperazine esters

INVENTOR(S): Kato, Hideo; Nishikawa, Tomoyasu; Mouri, Takaaki

PATENT ASSIGNEE(S): Hokuriku Pharmaceutical Co., Ltd.

SOURCE: Ger. Offen., 15 pp.

CODEN: GWXXBX

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE | | |
|------------------------|------|----------|-----------------|----------|--|--|
| | | | | | | |
| DE 2355420 | A1 | 19740704 | DE 1973-2355420 | 19731106 | | |
| JP 49069683 | Α | 19740705 | JP 1972-111219 | 19721108 | | |
| JP 49132088 | Α | 19741218 | | | | |
| JP 52010877 | В | 19770326 | | | | |
| PRIORITY APPLN. INFO.: | | | US 1972-254403 | 19720518 | | |

GI For diagram(s), see printed CA Issue.

AB (Homo)piperazinylalkyl [m, n = 2, 3; R = e.g. C6H2-(OMe)3-2,3,4, C6H4Cl-2, C6H4F-4, Ph, 2-, 3-, or 4-pyridyl, CH:-CHPh, CH:CHC6H2(OMe)3-2,3,4] and their salts were prepd.by refluxing 1-(hydroxyalkyl)(homo)piperazines and RCO2H inC6H6 in the presence of p-MeC6H4SO3H with H2O removal. I were useful as analgesics, anticonvulsants, antihypertensives, and coronary blood vessel dilators, especially in the treatment of circulatory diseases (no data).

IT 90-50-6 33130-03-9

RN 90-50-6 CAPLUS

CN 2-Propenoic acid, 3-(3,4,5-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

RN 33130-03-9 CAPLUS

CN 2-Propenoic acid, 3-(2,3,4-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

MeO CH
$$=$$
 CH $-$ CO₂H

L9 ANSWER 32 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1999:455150 CAPLUS

DOCUMENT NUMBER: 131:252293

TITLE: Novel Hypotensive Agents from Verbesina caracasana. 6.

> Synthesis and Pharmacology of Caracasandiamidel Carmignani, Marco; Volpe, Anna R.; Delle Monache,

AUTHOR (S): Franco; Botta, Bruno; Espinal, Romulo; De Bonnevaux, Stella C.; De Luca, Carlo; Botta, Maurizio; Corelli, Federico; Tafi, Andrea; Ripanti, Giuseppe; Delle

Monache, Giuliano

CORPORATE SOURCE: Dipartimento di Biologia di Base e Applicata Sezione

di Farmacologia, Universita di L'Aquila, Coppito,

67010, Italy

SOURCE: Journal of Medicinal Chemistry (1999), 42(16),

3116-3125

CODEN: JMCMAR; ISSN: 0022-2623

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

Caracasandiamide, a second hypotensive agent isolated from Verbesina caracasana, is the cyclobutane dimer (truxinic type) of the previously reported 1-[(3,4-dimethoxycinnamoyl)amino]-4-[(3-methyl-2butenyl)guanidino]butane (caracasanamide) (Delle Monache, G.; et al. BioMed. Chemical Lett. 1992, 25, 415-418). The structure was confirmed by synthesis starting from β -truxinic acid obtained by photoaddn. of 3,4-dimethoxycinnamic acid. The dimer was coupled with 2 mol of prenylagmatine to give caracasandiamide in satisfactory yield. By contrast, the direct photodimerization of caracasanamide was unsuccessful. Caracasandiamide, assayed by the iv route in anesthetized rats at doses ranging from 50 to 3200 $\mu g/kg$ of body weight, was found to have no appreciable effect on heart rate. At lower doses, the drug stimulates breathing and increases cardiac inotropism, stroke volume, and cardiac output, thus augmenting blood pressure and aortic flow. At higher doses, caracasandiamide depresses breathing likely through central neurogenic mechanisms (not involved in the cardiovascular effects), continues to stimulate cardiac inotropism, and induces, by reducing peripheral vascular resistance, arterial hypotension with reduction of both aortic flow and stroke volume These cardiovascular effects appear to involve complex interactions at the level of the peripheral $\beta1$ -, $\beta2$ -, and $\alpha 2$ -adrenoreceptor-dependent as well as M2- and M4-cholinergic receptor-dependent transductional pathways both in cardiovascular myocells and at the level of the postganglionic sympathetic endings (with reserpine- and guanethidine-like mechanisms). The cardiovascular effects of caracasandiamide, different from those of caracasanamide, do not depend on significant actions on the central nervous system and on baroreflex pathways. In a similar manner and more effective than caracasanamide, caracasandiamide may be considered a hypotensive and antihypertensive drug. It is devoid of some of the neg. side effects, e.g., reflex tachycardia and decreased cardiac inotropism, which are shown by the majority of the most common antihypertensive and vasodilator drugs.

IT 146269-40-1, Z-Caracasanamide RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(cardiovascular effects of caracasandiamide: comparison with other antihypertensives and vasodilators)

146269-40-1 CAPLUS ŔŊ

2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[[imino[(3-methyl-2-CNbutenyl)amino]methyl]amino]butyl]-, (2Z)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

$$\begin{array}{c|c} & H & H & H \\ \hline Z & & N & \\ \hline & & (CH_2) \stackrel{H}{4} & N & \\ \hline & & NH & \\ \hline & & \\ & &$$

REFERENCE COUNT: 40 THERE ARE 40 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 CAPLUS COPYRIGHT 2007 ACS on STN ANSWER 33 OF 98

ACCESSION NUMBER: 1998:698915 CAPLUS

DOCUMENT NUMBER: 130:60839

TITLE: Inhibitory effect of tranilast on hypertrophic

collagen production in the spontaneously

hypertensive rat heart

AUTHOR (S): Umemura, Kazuo; Kikuchi, Shinji; Suzuki, Yasuhiro;

Nakashima, Mitsuyoshi

Department of Pharmacology, Hamamatsu University CORPORATE SOURCE:

School of Medicine, Hamamatsu, 431 - 31, Japan

SOURCE: Japanese Journal of Pharmacology (1998), 78(2),

161-167

CODEN: JJPAAZ; ISSN: 0021-5198 Japanese Pharmacological Society

DOCUMENT TYPE: Journal LANGUAGE: English

PUBLISHER:

not

TΤ

Tranilast, N-(3,4-dimethoxycinnamoyl)anthranilic acid, a widely used antiallergy drug in Japan, has been shown to inhibit transforming growth factor-β1 release from fibroblasts and reduce collagen synthesis in keloid cells. In the present study, we have investigated the effect of this drug on cardiac hypertrophy in spontaneously hypertensive rats (SHR), with a focus on the cardiac collagen matrix, which is associated with myocardial stiffness. Twenty-four-week-old SHRs and Wistar Kyoto rats (WKYs) were administered translast (300 mg/kg) orally once a day for 4 wk. This treatment significantly suppressed increases in left ventricular collagen concentration (P<0.05) and the left ventricular weight/body

wts. ratios (P<0.05) in SHRs, and tranilast was ineffective on collagen concentration and ventricular weight/body wts. ratios in WKYs. Tranilast did

affect systolic or diastolic blood pressure, end-diastolic left ventricular pressure and heart rate in both SHRs and WKYs, and the agent did not change pos. dp/dt or cardiac output in SHRs. The pressure-volume relation curve was shifted to the left by the drug; the slope (k) of the logarithm of the pressure-volume relation curve was significantly increased (P<0.05) in SHRs. It is concluded that the suppression of increases in cardiac collagen and left ventricular mass by tranilast results in a corresponding prevention of cardiac stiffness as studied in the SHR.

53902-12-8, Tranilast RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(inhibitory effect of tranilast on hypertrophic collagen production in the spontaneously hypertensive rat heart)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 34 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2005:327442 CAPLUS

DOCUMENT NUMBER:

142:456667

TITLE:

Effect of Sodium Ferulate on Migration of Vascular

Smooth Muscle Cells Induced by Platelet Derived Growth

Factor and Endothelin-1

AUTHOR (S):

Han, Ying; Xie, Liangdi; Xu, Changsheng; Wang, Huajun

CORPORATE SOURCE: The First Affiliated Hospital, Fujian Medical

University, Fuzhou, Fujian Province, 350005, Peop.

Rep. China

SOURCE:

Zhongguo Dongmai Yinghua Zazhi (2004), 12(6), 659-661

CODEN: ZDYZFM; ISSN: 1007-3949

PUBLISHER:

Zhongguo Dongmai Yinghua Zazhi Bianjibu

DOCUMENT TYPE: Journal LANGUAGE: Chinese

To investigate the effect of sodium ferulate (SF), one of the principal components of rhizoma ligustici wallichii, on the migration of vascular smooth muscle cells (VSMC) induced by endothelin-1 (ET-1) and platelet derived growth factor (PDGF), VSMC derived from spontaneously hypertensive rats (SHR) were cultured. Cell migration was determined by modified Boyden chamber assays. [Ca2+]i was measured with fluorescent Ca2+ indicator Fura-2/AM. The results showed that ET-1 and PDGF significantly induced a migration of VSMC in a dose-dependent manner, which was inhibited by pretreatment of VSMC with SF (107-10-3 mol/L) dose-dependence. The peak inhibition rates of migration induced by ET-1 and PDGF were 85.04% and 81.92% resp. ET-1 and PDGF provoked the rise of [Ca2+]i in VSMC, which was significantly suppressed by 10-3 mol/L SF with a inhibitory peak at 80.14% and 76.69%. The cell migration and rise of [Ca2+]i induced by ET-1 and PDGF in VSMC from SHR may be suppressed by SF. IT 24276-84-4, Sodium ferulate

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(effect of sodium ferulate on migration of vascular smooth muscle cells induced by platelet derived growth factor and endothelin-1)

RN 24276-84-4 CAPLUS

CN 2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)-, monosodium salt (9CI) (CA INDEX NAME)

$$CH = CH - CO_2H$$

OMe

Na

L9 ANSWER 35 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1982:582873 CAPLUS

DOCUMENT NUMBER: 97:182873

TITLE: Bicyclic compounds and their use

INVENTOR(S): Oka, Yoshikazu; Nishikawa, Kohei; Miyake, Akio

PATENT ASSIGNEE(S): Takeda Chemical Industries, Ltd., Japan

SOURCE: Eur. Pat. Appl., 52 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PA' | TENT NO. | | | KINI |) | DATE | | AP | PLICATION NO. | DATE |
|-----|----------|-----|-----|--------|----------|-----------|-----|----|---------------|----------|
| | 51391 | | | A1 | | 19820512 | | ΕP | 1981-304940 | 19811021 |
| ΕP | 51391 | | | B1 | | 19840905 | | | | |
| | R: AT, | BE, | CH, | DE, | FR | , IT, LU, | NL, | S | 3 | |
| JP | 57077651 | | | Α | | 19820515 | | JP | 1980-154394 | 19801031 |
| JP | 62012800 | | | В | | 19870320 | | | | |
| JP | 57179141 | | | A | | 19821104 | | JP | 1981-64371 | 19810428 |
| JP | 02023538 | | | В | | 19900524 | | | | |
| ΑU | 8176521 | | | Α | | 19820506 | | ΑU | 1981-76521 | 19811016 |
| | 543804 | | | B2 | | 19850502 | | | | |
| | 4822818 | | | Α | | 19830524 | | | 1981-312639 | 19811019 |
| | 8107253 | | | Α | | 19820929 | | ZA | 1981-7253 | 19811020 |
| | 2086393 | | | Α | | 19820512 | | GB | 1981-31719 | 19811021 |
| | 2086393 | | | В | | 19840111 | | | | |
| | 9220 | | | T | | 19840915 | | | 1981-304940 | 19811021 |
| | 8103383 | | | Α | | 19820501 | | FΙ | 1981-3383 | 19811028 |
| | 73698 | | | В | | 19870731 | | | | |
| | 73698 | | | С | | 19871109 | | | , | |
| | 8104781 | | | Α | | 19820501 | | DK | 1981-4781 | 19811029 |
| | 164917 | | | В | | 19920907 | | | | |
| | 164917 | | | C | | 19930201 | | | | |
| | 8103662 | | | Α | | 19820503 | | ИО | 1981-3662 | 19811029 |
| | 155133 | | | В | | 19861110 | | | | |
| | 155133 | | | С | | 19870218 | | | | |
| | 28805 | | | A2 | | 19831228 | | HU | 1981-3176 | 19811029 |
| | 183652 | | | В | | 19840528 | | | | |
| | 1271372 | | | A3 | | 19861115 | | | 1981-3350151 | 19811029 |
| | 1287444 | | | С | | 19910806 | | | 1981-389042 | 19811029 |
| | 506714 | | | A1 | | 19830601 | | | 1981-506714 | 19811030 |
| | 515269 | | | A1 | | 19831201 | | | 1982-515269 | 19820826 |
| | 4474692 | | | A | | 19841002 | | | 1983-494061 | 19830512 |
| | 524148 | | | A1 | | 19850501 | | | 1983-524148 | 19830715 |
| | 1287446 | | | C2 | | 19910806 | | - | 1984-468185 | 19841119 |
| | 8602859 | | | A | | 19820503 | | ИО | 1986-2859 | 19860715 |
| | 157103 | | | В | | 19871012 | | | | |
| | 157103 | | | C A | | 19880120 | | TD | 1005 2000 | 10050010 |
| υP | 63002963 | | | A | | 19880107 | | υP | 1987-30000 | 19870212 |

| JP 02024265 | В | 1990 | 0529 | | | | |
|------------------------|---|------|------|----|-------------|----|----------|
| US 5098892 | Α | 1992 | 0324 | US | 1989-302940 | | 19890130 |
| PRIORITY APPLN. INFO.: | | | | JР | 1980-154394 | Α | 19801031 |
| | | | | JР | 1981-64371 | Α | 19810428 |
| | | | | US | 1981-312639 | A3 | 19811019 |
| | | | | ΕP | 1981-304940 | Α | 19811021 |
| | | | | CA | 1981-389042 | A | 19811029 |
| | | | | NO | 1981-3662 | A1 | 19811029 |
| | | | | | | | |

OTHER SOURCE(S): GI

CASREACT 97:182873

$$(C_nH_{2n-1})$$

$$NCOCHR^3NHCHR^4COR^5$$

$$(CH_2)_mCO_2R^2$$

AB Peptide derivs. I [R, R1 = H, OH, C1-4 alkoxy; RR1 = C1-4 alkylenedioxy; R2 = H, C1-4 alkyl; R3 = H, C1-4 alkyl, (un) substituted amino-C1-4 alkyl; R4 = H, C1-4 alkyl, (un) substituted phenyl-C1-4 alkyl; R5 = OH, C1-4 alkoxy, mono- or di-C1-4 alkylamino; m, n = 1, 2] were prepared as angiotensin-converting enzyme (ACE) inhibitors and antihypertensives. Thus, H-Gly-OEt.HCl was treated with 2-indanone in MeOH containing NaBH3CN to give indanylglycine II, which was coupled with PhCH2O2C-Ala-OH by ClCO2CH2CHMe2 to give the protected dipeptide, which was deblocked by saponification and hydrogenolysis to give dipeptide III (R6 = H). The latter was treated with PhCH2CH2COCO2Et in EtOH for 1 h at room temperature and the resulting solution was reduced by NaBH3CN

and then treated with HCl/EtOH to give III.HCl [R = PhCH2CH2CH(CO2Et)] (IV). IV at 1 μ M inhibited ACE by 87%.

IT 2316-26-9P

> RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation and esterification of, with ethanol)

RN 2316-26-9 CAPLUS

CN 2-Propenoic acid, 3-(3,4-dimethoxyphenyl)- (9CI) (CA INDEX NAME)

ANSWER 36 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1983:107786 CAPLUS

DOCUMENT NUMBER:

98:107786

TITLE:

Alicyclic compounds and their use

INVENTOR(S):

Oka, Yoshikazu; Nishikawa, Kohei; Miyake, Akio

Takeda Chemical Industries, Ltd., Japan

SOURCE:

Eur. Pat. Appl., 39 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

English

2 .

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT ASSIGNEE(S):

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|----------|----------------------|--------------------|----------|
| EP 57998 EP 57998 | A1 B1 | 19820818 19840808 | EP 1982-300300 | 19820121 |
| R: BE, CH, DE, | FR, GB | , IT, NL, SE | | |
| JP 57123151 | A | 19820731 | JP 1981-9470 | 19810123 |
| JP 57203050 | Α | 19821213 | JP 1981-88539 | 19810609 |
| JP 58109458 | A | 19830629 | JP 1981-208817 | 19811222 |
| PRIORITY APPLN. INFO.: | | | JP 1981-9470 A | 19810123 |
| | | | JP 1981-88539 A | 19810609 |
| • | | | JP 1981-208817 A | 19811222 |
| OTHER SOURCE(S): | CASREA | CT 98:107786 | ; MARPAT 98:107786 | |

GI

ΔR Peptide derivs. CylN(CH2CO2R)COCHR1NHCHR2CO2R3 [Cyl = C3-10 alicyclic group; R = H, alkyl, R1 = H, alkyl, aralkyl; R2 = H, alkyl, (un) substituted aralkyl; R3 = H, alkyl] were prepared as antihypertensives due to their ability to inhibit angiotensin-converting enzyme (ACE). Thus, H-Gly-OEt.HCl underwent reductive cycloalkylation with cyclopentanone in the presence of NaBH3CN to give cyclopentylglycine I, which was coupled with Z-Ala-OH (Z = PhCH2O2C) by ClCO2CH2CHMe2 in THF containing Et3N to give peptide II (R4 = Z, R5 = Et), which was saponified and then Z-deblocked by hydrogenolysis to give II (R4 = R5 = H). The latter was treated with PhCH2CH2COCO2Et in the presence of NaBH3CN to give peptide derivative III (as the HCl salt). III at 1 μM inhibited ACE by 84%.

IT 2316-26-9P

> RL: SPN (Synthetic preparation); PREP (Preparation) (preparation and purification of, with ethanol)

RN 2316-26-9 CAPLUS

CN2-Propenoic acid, 3-(3,4-dimethoxyphenyl)- (9CI) (CA INDEX NAME)

ANSWER 37 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN T.9

ACCESSION NUMBER: 1999:769090 CAPLUS

132:87936 DOCUMENT NUMBER:

Novel hypotensive agents from Verbesina caracasana. 7. TITLE:

Further hypotensive metabolites from verbesina

AUTHOR(S): Delle Monache, Giuliano; Volpe, Anna Rita; Delle

Monache, Franco; Vitali, Alberto; Botta, Bruno; Espinal, Romulo; De Bonnevaux, Stella C.; De Luca, Carlo; Botta, Maurizio; Corelli, Federico; Carmignani,

Marco

Centro Chimica dei Recettori, Centro Chimica dei CORPORATE SOURCE:

Recettori, Universita Cattolica, Rome, 00168, Italy

Bioorganic & Medicinal Chemistry Letters (1999), SOURCE:

9(22), 3249-3254

CODEN: BMCLE8; ISSN: 0960-894X

PUBLISHER: Elsevier Science Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

AB After the isolation of caracasanamide and caracasandiamide, further hypotensive components of Verbesina caracasana were shown to be N3-prenylagmatine, N1-3',4'-dimethoxycinnamoylagmatine, agmatine and galegin (prenylguanidine). The structures were assigned on the basis of the spectral data of both metabolites and products from their alkaline hydrolyzes. A pharmacol. anal. of these products is also presented.

IT 128009-18-7P

> RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(isolation of antihypertensive guanidine metabolites from

Verbesina caracasana)

128009-18-7 RNCAPLUS

CN 2-Propenamide, N-[4-[(aminoiminomethyl)amino]butyl]-3-(3,4dimethoxyphenyl) - (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{O} \\ \parallel \\ \text{CH} \end{array} \begin{array}{c} \text{CH} \longrightarrow \text{CH} - \text{CH} - \text{NH} - \text{CH}_2)_4 - \text{NH} - \text{CH}_2 \\ \text{MeO} \end{array}$$

REFERENCE COUNT: 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 38 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1991:429185 CAPLUS

DOCUMENT NUMBER:

TITLE:

115:29185

2-Aryl-substituted benzannulated 5-ring heterocycles

as potential cardiovascular agents. 1

AUTHOR (S):

Rose, Ulrich

CORPORATE SOURCE:

Inst. Pharm., Johannes Gutenberg-Univ., Mainz, D-6500,

SOURCE:

Chemiker-Zeitung (1991), 115(2), 55-8

CODEN: CMKZAT; ISSN: 0009-2894

DOCUMENT TYPE:

Journal

LANGUAGE:

German

GI

AB Benzazoles I (X = S, O; R = substituted Ph, pyridyl, styryl) were prepared from RCO2H and 2-HXC6H4NH2. I (X = O, R = 2-methylthio-3-pyridyl) had 48% of the antihypertensive activity of fostedil. I also have fungicidal activity.

IT 90-50-6

RL: RCT (Reactant); RACT (Reactant or reagent)

(cyclization of, with aminothiophenol and aminophenol)

RN 90-50-6 CAPLUS

CN 2-Propenoic acid, 3-(3,4,5-trimethoxyphenyl) - (9CI) (CA INDEX NAME)

L9 ANSWER 39 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1978:105114 CAPLUS

DOCUMENT NUMBER: 88:105114

TITLE: 2-[(3',5'-Dimethoxy-4'-hydroxy)phenylmethylene]-3-

carboxy-4-(3",5"-dimethoxy-4"-hydroxyphenyl)- γ -

butyrolactone

INVENTOR(S): Umezawa, Hamao; Takeuchi, Tomio; Kumada, Yoshiki

PATENT ASSIGNEE(S): Microbiochemical Research Foundation, Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 6 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|------------------------|------|----------|-----------------|---|----------|
| JP 52136163 | Α | 19771114 | JP 1976-53527 | | 19760510 |
| PRIORITY APPLN. INFO.: | | | JP 1976-53527 | A | 19760510 |

$$MeO$$
 HO_2C OMe OM

and air introduced vigorously over 3 h to give 45.1% the title compound (I). I at 50 mg/kg orally decreased the blood pressure of spontaneously hypertensive rats by 23.1% in 6 h.

IT 530-59-6

RL: RCT (Reactant); RACT (Reactant or reagent)

(self-cyclocondensation of)

RN 530-59-6 CAPLUS

CN 2-Propenoic acid, 3-(4-hydroxy-3,5-dimethoxyphenyl)- (9CI) (CA INDEX NAME)

L9 ANSWER 40 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2003:43029 CAPLUS

DOCUMENT NUMBER: 138:106536

TITLE: Preparation of substituted dicinnamoylquinides and

their therapeutic use in augmentation of adenosine

function

INVENTOR(S): De Paulis, Tomas; Lovinger, David M.; Martin, Peter R.

PATENT ASSIGNEE(S): Vanderbilt University, USA SOURCE: U.S. Pat. Appl. Publ., 17 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|-----------------------------------|----------|------------|------------------------------------|---|----------------------|
| US 2003013758 US 6693128 | A1 B2 | 20030116 | US 2002-143606 | | 20020510 |
| CA 2347879 PRIORITY APPLN. INFO.: | A1 | 20021111 | CA 2001-2347879 US 2001-290282P | p | 20010516 20010511 |
| OTHER SOURCE(S): | MADDAT | 138:106536 | CA 2001-2347879 | A | 20010511 |
| GI | MARPAT | 138:106536 | | | |

AB This invention describes the preparation of dicinnamoylquinides, such as I [R = H; R1-6 = H, OH, alkyl, alkoxy, halogen], and their use as therapeutic agents for enhancing adenosine levels in the brain and peripheral organs.

These agents, which partially or completely inhibit adenosine transport, are particularly useful in treating human diseases or conditions that benefit from acute or chronic elevated levels of adenosine, such as reperfusion injury, coronary or cerebral ischemia, coronary vasoconstriction, paroxysmal supraventricular tachycardia, hypertension, wound healing, diabetes, inflammation, stroke, depression, cardiovascular disorders, or sleep disturbances. quinides can also be used to protect normal cells from chemotoxicity in patients undergoing cancer therapy and reverse the behavioral effects of caffeine intake. Thus, O-protected (-)-quinic acid γ -lactone II (R = CO2CH2CCl3) was acylated with the in situ formed acid chloride of 4-chlorocinnamic acid to give the 3,4-di-(4-chlorocinnamoy1)-1,5-quinide I (R = CO2CH2CCl3, R1 = R3 = R4 = R6 = H, R2 = R5 = Cl) with 79% yield. Subsequent deprotection of the quinide using zinc powder and AcOH in THF gave the desired quinide I (R = R1 = R3 = R4 = R6 = H, R2 = R5 = C1) in 78% yield. The prepared quinides were evaluated for adenosine transporter affinity and for inhibition of adenosine transport. Also, pharmaceutical compns. containing these quinides were presented.

IT 331-39-5, 3,4-Dihydroxycinnamic acid 1135-24-6, 3-Methoxy-4-hydroxycinnamic acid

RL: RCT (Reactant); RACT (Reactant or reagent)
(preparation of substituted dicinnamoylquinides and their therapeutic use in augmentation of adenosine function)

RN 331-39-5 CAPLUS

CN 2-Propenoic acid, 3-(3,4-dihydroxyphenyl)- (9CI) (CA INDEX NAME)

$$_{
m HO}$$
 $_{
m OH}$ $_{
m CH}$ $_{
m CH}$ $_{
m CH}$ $_{
m CH}$

RN 1135-24-6 CAPLUS

CN 2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)- (9CI) (CA INDEX NAME)

$$CH = CH - CO_2H$$

IT 485402-21-9P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of substituted dicinnamoylquinides and their therapeutic use in augmentation of adenosine function)

RN 485402-21-9 CAPLUS

CN 2-Propenoic acid, 3-[3-methoxy-4-[[(2,2,2-trichloroethoxy)carbonyl]oxy]phe nyl]-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L9 ANSWER 41 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1996:58393 CAPLUS

DOCUMENT NUMBER: 124:232440

TITLE: Thazolidinedione compounds useful as antidiabetics

INVENTOR(S): Regnier, Gilbert; Charton, Yves; Duhault, Jacques;

Espinal, Joseph

PATENT ASSIGNEE(S): Adir et Compagnie, Fr.

SOURCE: U.S., 7 pp. Cont.-in-part of U.S. Ser. No. 133,898,

abandoned.

CODEN: USXXAM PATENT TYPE: Patent

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--------|------------|-------------------|----------|
| | | | | |
| US 5478853 | A | 19951226 | US 1995-376052 | 19950120 |
| FR 2696743 | A1 | 19940415 | FR 1992-12123 | 19921012 |
| FR 2696743 | B1 | 19941223 | | |
| PRIORITY APPLN. INFO.: | | | FR 1992-12123 A | 19921012 |
| | | | US 1993-133898 B2 | 19931012 |
| OTHER SOURCE(S): | MARPAT | 124:232440 | | |

GI

$$Ar-A-X-B$$
 CH_2
 NH
 S

$$Q^{1}=$$
C1
CONHCH₂CH₂
OMe

The title compds. are 5-(4-substituted benzyl)thiazolidine-2,4-diones I [Ar = polymethylene ring with optional alkyl substituent(s), (un)substituted aryl or heterocyclyl; A = bond, hydrocarbondiyl with double bond, (CH2)1-3, CMe2(CH2)0-2, (un)substituted CHPh(CH2)0-2, O(CH2)1-3, S(CH2)1-3; X = O, CONR, SO2NR; R = H, alkyl, alkenyl; or ArAX = phthalimido; B = saturated hydrocarbondiyl with optional OH or oxo substituent] and their enantiomers, diastereoisomers, and pharmaceutically tolerable salts. The compds. are useful for treating insulin resistance

and/or non-insulin-dependent diabetes, possibly associated with hypertension. An exemplary compound compound is 5-[4-[2-(2-methoxy-5-chlorobenzamido)ethyl]benzyl]thiazolidine-2,4-dione, i.e., I [Ar-A-X-B- = Q1] (II), which was prepared by cyclization of the corresponding 3-phenyl-2-chloropropionic acid derivative with thiourea in sulfolane at 120°, followed by hydrolysis with aqueous HCl at 100°. II, at \leq 10 mg/kg/day orally in mice, had the same hypoglycemic effect as ciglitazone at 50-100 mg/kg/day. II also had little or no hematol. effect at 250 mg/kg/day in rats, whereas pioglitazone had strong adverse effects at 100 mg/kg/day.

IT 174772-41-9P 174772-45-3P 174772-47-5P
 RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of thiazolidinediones as antidiabetics)

RN 174772-41-9 CAPLUS

CN

2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[2-[4-[(2,4-dioxo-5-thiazolidinyl)methyl]phenyl]ethyl]- (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 2-A

PAGE 1-A

PAGE 2-A

174772-47-5 CAPLUS RN

2-Propenamide, 3-[4-(acetyloxy)-3-methoxyphenyl]-N-[2-[4-[(2,4-dioxo-5-thiazolidinyl)methyl]phenyl]ethyl]- (9CI) (CA INDEX NAME) CN

PAGE 2-A

ANSWER 42 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN L9

ACCESSION NUMBER:

PATENT ASSIGNEE(S):

1977:139646 CAPLUS

DOCUMENT NUMBER:

86:139646

TITLE:

Vasoactive N-benzoyl- or N-cinnamoylglycines

Zambeletti Espana S. A., Spain

SOURCE:

DOCUMENT TYPE:

Span., 9 pp. CODEN: SPXXAD

LANGUAGE:

Patent

FAMILY ACC. NUM. COUNT:

Spanish

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE -------------------ES 422191 **A**1 19760416 ES 1974-422191 19740110 PRIORITY APPLN. INFO.: A 19740110 ES 1974-422191 GI

AB N-acylglycines I and II (R = H, HO, AcO, MeO, Cl; R1 = H, HO, AcO; R2 = H, Et) were prepared by acylation of glycine or its Et ester with the appropriate benzoyl or cinnamoyl chloride. Some I and II had hypertensive and others hypotensive activity in tests on the cat (in vivo) and the rabbit (in vitro, hypertension induced with noradrenaline). Thus, H2NCH2CO2Et 20.4, 4,3,5-(AcO)(MeO)2C6H2COCl 38 g, and NaHCO3 112 g in AcOEt-Et2O-H2O was stirred 3 h at room temperature and 4 h at 40°C to give 15 g I (R = MeO, R1 = AcO, R2 = Et), which was hydrolyzed to the acid.

IT 62098-72-0P 62098-77-5P

RL: SPN (Synthetic preparation); PREP (Preparation) (preparation and effect on blood pressure)

RN 62098-72-0 CAPLUS

CN Glycine, N-[3-[4-(acetyloxy)-3,5-dimethoxyphenyl]-1-oxo-2-propenyl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 62098-77-5 CAPLUS CN Glycine, N-[3-(4-hydroxy-3,5-dimethoxyphenyl)-1-oxo-2-propenyl]- (9CI) (CA INDEX NAME)

L9 ANSWER 43 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2000:843218 CAPLUS

DOCUMENT NUMBER:

133:350051

TITLE:

Preparation process of [3-(4-Hydroxy-3-

methoxyphenyl)propenamido]-N-ethyl nitrate having

cardiovascular pharmacological activity

INVENTOR(S):

Xu, Jingfeng; Wang, Jinping; Qi, Ping; Yang, Yongge;

Zhang, Mei

PATENT ASSIGNEE(S):

Peop. Rep. China

SOURCE:

Faming Zhuanli Shenqing Gongkai Shuomingshu, 16 pp.

CODEN: CNXXEV

DOCUMENT TYPE:

Patent

LANGUAGE:

Chinese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| KIND | DATE | APPLICATION NO. | DATE |
|------|----------|-----------------|---|
| | | | |
| Α | 20000405 | CN 1999-119659 | 19990924 |
| В | 20020501 | | |
| | | CN 1999-119659 | 19990924 |
| | A | A 20000405 | A 20000405 CN 1999-119659 B 20020501 |

OTHER SOURCE(S): CASREACT 133:350051

Title compound was prepared by chlorinating 3-(4-acetoxy-3-methoxyphenyl)propanoic acid with oxalyl chloride or SOC12 in dichloromethane in the presence of DMF at 15°-80° for 1-5 h to obtain 3-(4-acetoxy-3-methoxyphenyl)propanoyl chloride, acylating with 2-aminoethyl nitrate in toluene-water to obtain [3-(4-acetoxy-3-methoxyphenyl)propenamido]-N-Et nitrate as intermediate. Title compound was tested for cardiovascular pharmacol. activity.

IT 306272-46-8P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation process of hydroxymethoxyphenylpropenamidoethyl nitrate having cardiovascular pharmacol. activity)

RN 306272-46-8 CAPLUS

CN 2-Propenamide, 3-(4-hydroxy-3-methoxyphenyl)-N-[2-(nitrooxy)ethyl]- (9CI) (CA INDEX NAME)

IT 2596-47-6, 3-(4-Acetoxy-3-methoxyphenyl)prop-2-enoic acid

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation process of hydroxymethoxyphenylpropenamidoethyl nitrate having cardiovascular pharmacol. activity)

RN 2596-47-6 CAPLUS

CN 2-Propenoic acid, 3-[4-(acetyloxy)-3-methoxyphenyl]- (9CI) (CA INDEX NAME)

IT 306272-47-9P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation process of hydroxymethoxyphenylpropenamidoethyl nitrate having cardiovascular pharmacol. activity)

RN 306272-47-9 CAPLUS

CN 2-Propenamide, 3-[4-(acetyloxy)-3-methoxyphenyl]-N-[2-(nitrooxy)ethyl](9CI) (CA INDEX NAME)

L9 ANSWER 44 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1997:791311 CAPLUS

DOCUMENT NUMBER: 128:123446

TITLE: A multidisciplinary research on Verbesina caracasana

AUTHOR(S): Botta, Bruno; Misiti, Domenico; Delle Monache,

Giuliano; Persichilli, Silvia; Vitali, Alberto; Botta,

Maurizio; Corelli, Federico; Carmignani, Marco

CORPORATE SOURCE: Dipartimento di Studi di Chimica e Tecnologia delle

Sostanze Biologicamente Attive, Universita "La

Sapienza", Rome, I-00185, Italy

SOURCE: Gazzetta Chimica Italiana (1997), 127(6), 305-310

CODEN: GCITA9; ISSN: 0016-5603

PUBLISHER: Societa Chimica Italiana

DOCUMENT TYPE: Journal LANGUAGE: English

AB Two hypotensive agents have been isolated from Verbesina caracasana Fries and attributed the structure of 1-[(3,4-dimethoxycinnamoyl)amino]-4-[(3-methyl-2-butenyl)guanido]butane and β -truxinic[bis-3',4'-dimethoxy]di[N-(3-methylbut-3-enyl)guanidobutyl]amide, resp. The structures of the two compds. have been confirmed by syntheses and their pharmacol. profiles established. Studies on cell cultures of Verbesina caracasana gave unexpected results showing that a β -glucosyltransferase is active in whole cells.

IT 146269-39-8P 146269-40-1P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PUR (Purification or recovery); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(isolation, preparation and pharmacol. of Verbesina caracasana constituents)

RN 146269-39-8 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[[imino[(3-methyl-2-butenyl)amino]methyl]amino]butyl]-, (E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

$$\begin{array}{c|c} & & & \\ &$$

RN 146269-40-1 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[[imino[(3-methyl-2-butenyl)amino]methyl]amino]butyl]-, (2Z)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

$$\begin{array}{c|c} & H & H & H \\ \hline Z & O & NH & N \\ \hline \\ O & O & NH & N \\ \hline \end{array}$$

IT 130973-10-3

RL: RCT (Reactant); RACT (Reactant or reagent)

(isolation, preparation and pharmacol. of Verbesina caracasana constituents)

RN 130973-10-3 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-, (E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

REFERENCE COUNT:

11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 45 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1985:197969 CAPLUS

DOCUMENT NUMBER:

102:197969

TITLE:

Validity of the Oriental medicines. Part 74. Liver

protective drugs. Part 20. Studies on the

constituents of Ephedra. Part 17. Pharmacological actions of analogs of feruloylhistamine, an imidazole

alkaloid of Ephedra roots

AUTHOR(S):

Hikino, Hiroshi; Kiso, Yoshinobu; Ogata, Minoru; Konno, Chohachi; Aisaka, Kazuo; Kubota, Hiroaki;

Hirose, Nakako; Ishihara, Takafumi

CORPORATE SOURCE:

Pharm. Inst., Tohoku Univ., Sendai, Japan

SOURCE:

Planta Medica (1984), 50(6), 478-80

CODEN: PLMEAA; ISSN: 0032-0943

DOCUMENT TYPE:

Journal

LANGUAGE:

English

GI

MeO

$$CH = CHCONHCH_2CH_2$$
 N

- AB Analogs of feruloylhistamine (I), an imidazole alkaloid of Ephedra roots, were prepared and their pharmacol. actions determined These compds. had hypotensive, histidine decarboxylase [9024-61-7] inhibitory, antiulcer, and antihepatotoxic actions.
- IT 94848-18-7DP, analogs 94848-21-2P 94848-23-4P

RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses)

(preparation and pharmacol. of)

RN 94848-18-7 CAPLUS

CN 2-Propenamide, 3-(4-hydroxy-3-methoxyphenyl)-N-[2-(1H-imidazol-4-yl)ethyl](9CI) (CA INDEX NAME)

$$\begin{array}{c}
H \\
N \\
\end{array}$$

$$\begin{array}{c}
CH_2 - CH_2 - NH - C - CH = CH \\
\end{array}$$
OH
OMe

RN 94848-21-2 CAPLUS

CN 2-Propenamide, 3-(4-hydroxy-3,5-dimethoxyphenyl)-N-[2-(1H-imidazol-4-yl)ethyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c}
H & O \\
N & CH_2 - CH_2 - NH - C - CH - CH - CH - OMe
\end{array}$$
OMe

RN 94848-23-4 CAPLUS

CN 2-Propenamide, 3-(3,4-dihydroxyphenyl)-N-[2-(1H-imidazol-4-yl)ethyl]-(9CI) (CA INDEX NAME)

$$\begin{array}{c}
H \\
N \\
\end{array}$$

$$\begin{array}{c}
CH_2 - CH_2 - NH - C - CH = CH \\
\end{array}$$

$$\begin{array}{c}
O \\
H \\
OH
\end{array}$$

$$\begin{array}{c}
OH
\end{array}$$

L9 ANSWER 46 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1991:185027 CAPLUS

DOCUMENT NUMBER:

114:185027

TITLE:

Preparation of trisubstituted benzene compounds for

treating congestive heart failure

INVENTOR(S):

Hawkins, Lynn D.

PATENT ASSIGNEE(S):

Warner-Lambert Co., USA

SOURCE:

U.S., 15 pp. Cont. of U.S. Ser. No. 38,252, abandoned.

CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

1

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

US 4971959 A 19901120 US 1988-292580 19881230 US 5274002 A 19931228 US 1990-578965 19900906 PRIORITY APPLN. INFO.: US 1987-38252 B1 19870414 US 1988-292580 A3 19881230

OTHER SOURCE(S):

MARPAT 114:185027

GΙ

$$R^2X$$
 AY I
 R^2X
 R^2X
 R
 R
 R
 R
 R
 R

The title compds. [I; R1 = C3-6 cycloalkyl; R2 = alkyl; X = 0, S; A = bond, C1-7 alkylene, C2-6 alkenylene optionally interrupted by O, S, and imino; Y = CONR3R4 wherein R3, R4 = H, alkyl, azido, cyano] are prepared Hydrogenation of II [R = (E)-CH:CHCO2Me] (preparation given) over 5% Pd-C gave 83.9% propionate II (R = CH2CH2CO2Me), which was heated in anhydrous methanolic NH3 at 100° to give 32.0% I (R1 = cyclopentyl, R2X = MeO, A = CH2CH2, Y = CONH2) (III). III showed EC50 of 1 + 10-5M in improving coronary blood flow and increased heart contractility at 1.0 mg/kg in dogs.

IT 133332-31-7P

RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of, as cardiotonic and cardiovascular agent)

RN 133332-31-7 CAPLUS

CN 2-Propenoic acid, 3-[3-(cyclopentyloxy)-4-methoxyphenyl]-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L9 ANSWER 47 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1988:75410 CAPLUS

DOCUMENT NUMBER:

108:75410

TITLE:

Preparation of arylbenzothiazinones as calcium

antagonists

INVENTOR(S):

Lerch, Ulrich; Henning, Rainer; Kaiser, Joachim

Hoechst A.-G. , Fed. Rep. Ger.

PATENT ASSIGNEE(S): SOURCE:

Ger. Offen., 17 pp.

CODEN: GWXXBX

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------|------|----------|-----------------|----------|
| | | | | |
| DE 3614363 | A1 | 19871029 | DE 1986-3614363 | 19860428 |
| FI 8701813 | Α | 19871029 | FI 1987-1813 | 19870424 |
| EP 244723 | A2 | 19871111 | EP 1987-106067 | 19870425 |
| EP 244723 | A3 | 19881019 | | |

| R: AT, BE, CH | , DE, | ES, FR, GB, | GR, IT, LI, LU, NL, SI | Ξ | ÷ |
|------------------------|-------|--------------|------------------------|---|----------|
| AU 8771994 | Α | 19871029 | AU 1987-71994 | | 19870427 |
| AU 601659 | B2 | 19900913 | | | |
| DK 8702133 | Α | 19871029 | DK 1987-2133 | | 19870427 |
| NO 8701745 | A | 19871029 | NO 1987-1745 | | 19870427 |
| JP 62258371 | Α | 19871110 | JP 1987-102143 | | 19870427 |
| ZA 8702972 | Α | 19871125 | ZA 1987-2972 | | 19870427 |
| HU 45517 | A2 | 19880728 | HU 1987-1845 | | 19870427 |
| HU 199814 | В | 19900328 | | | |
| IL 82332 | Α | 19910816 | IL 1987-82332 | | 19870427 |
| PRIORITY APPLN. INFO.: | | | DE 1986-3614363 | Α | 19860428 |
| OTHER SOURCE(S): | CAS | REACT 108:75 | 410; MARPAT 108:75410 | | |
| GT | | | | | |

AΒ The title compds. [I; R1, R2, R3, R6, R7 = C1-4 alkyl, C1-3 alkoxy, F, C1, Br, CF3, NO2, OH, AcNH, NH2; R4 = H, C1-10 alkyl, C3-10 alkenyl, (substituted) Ph; R5 = H, C1-15 alkyl, C3-15 alkenyl, C4-8 cycloalkyl, cycloalkylalkyl, (substituted) Ph, phenylalkyl; R9, R10, R11, R12 = H, C1-10 alkyl, C1-6 alkanoyl, (substituted) phenylalkyl, benzhydryl, benzhydrylalkyl, phenylalkanoyl; A = (CH2)n X(CH2)n; X = CH2, O, S, CO, CHOH, CR82; R8 = H, C1-4 alkyl; B = NR9, piperazinyl, Q1, Q2, Q3, Q4; T =(CH2)p; D = CHOH, CO, NR13CO, O; R13, R14 = H, C1-4 alkyl; E = (CH2)q, CH:CH; F = bond, NR14CO, CO, O; G = aryl; m = 1-4; n = 1-3; p,q = 0-3] are prepared as Ca antagonists. A mixture of 3,4-dihydro-2-isopropyl-4-methyl-2-[2-(4-bromobutoxy)phenyl]-2H-1,4-benzothiazine-3-one, 3,4,5trimethoxyphenylacetate piperazide, K2CO3, and DMF was refluxed for 5 h to give 3,4-dihydro-2-isopropyl-4-methyl-2-[2-(4- (4-(2-(3,4,5trimethoxyphenyl)acetyl)piperazinyl)butoxy)phenyl]-2H-1,4-benzothiazine-3one.HCl. I displaced 3H-nitrendipine from membrane prepns. with IG50's of 10-6- 10-10 m.

IT 90-50-6, 3,4,5-Trimethoxycinnamic acid
RL: RCT (Reactant); RACT (Reactant or reagent)

(acylation by, of [(piperazinylbutoxy)phenyl]benzothiazinone)

RN 90-50-6 CAPLUS

CN 2-Propenoic acid, 3-(3,4,5-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

L9 ANSWER 48 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1975:453178 CAPLUS

DOCUMENT NUMBER: 83:53178

TITLE: Rigid amino acids related to α -methyldopa

AUTHOR(S): Cannon, Joseph G.; O'Donnell, John P.; Rosazza, John

P.; Hoppin, Charles R.

CORPORATE SOURCE: Coll. Pharm., Univ. Iowa, Iowa City, IA, USA

SOURCE: Journal of Medicinal Chemistry (1974), 17(5), 565-8

CODEN: JMCMAR; ISSN: 0022-2623

DOCUMENT TYPE: Journal LANGUAGE: English

GI For diagram(s), see printed CA Issue.

AB Title compds. (\pm) -2-amino-4,5-dihydroxyindan-2-carboxylic acid-HBr (I)

and (±)-2-amino-5,6-dihydroxy-1,2,3,4-tetrahydronaphthalene-2-

carboxylic acid-HBr (II), were prepared from the corresponding ketones by the Strecker reaction. I and II at concns. of 10-3M completely inhibited the decarboxylation of L-dopa by Dopa decarboxylase. The relation of activity to structure was discussed.

IT 331-39-5

RL: BIOL (Biological study)

(Dopa decarboxylase inhibition activity of, methyldopa analogs in

relation to)

RN 331-39-5 CAPLUS

CN 2-Propenoic acid, 3-(3,4-dihydroxyphenyl)- (9CI) (CA INDEX NAME)

L9 ANSWER 49 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2004:368857 CAPLUS

DOCUMENT NUMBER: 140:386000

TITLE: Compounds, compositions and methods for modulating fat

metabolism for treatment of metabolic disorders

INVENTOR(S): Gaudriault, Georges; Kilinc, Ahmet; Bousquet, Olivier;

Goupil-Lamy, Anne; Harosh, Itzik

PATENT ASSIGNEE(S): Obetherapy Biotechnology, Fr.

SOURCE: PCT Int. Appl., 461 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

WO 2004037159 A2 20040506 WO 2003-IL860 20031023
WO 2004037159 A3 20040715

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,

CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG AU 2003274652 **A1** 20040513 AU 2003-274652 20031023 PRIORITY APPLN. INFO.: US 2002-420316P P 20021023 WO 2003-IL860 W 20031023

OTHER SOURCE(S): MARPAT 140:386000

AB Methods and compns. of identifying candidate compds., for modulating fat metabolism and/or inhibiting Apobec-1 activity are provided. The invention relates to compds. and pharmaceutical compns. which are useful for regulating fat metabolism and can be used for treatment of diseases and disorders selected from the group consisting of overweight, obesity, atherosclerosis, hypertension, non-insulin dependent diabetes mellitus, pancreatitis, hypercholesteremia, hypertriglyceridemia, hyperlipidemia.

IT 686300-42-5

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(compds., compns. and methods for modulating fat metabolism for treatment of metabolic disorders)

RN 686300-42-5 CAPLUS

CN 2-Propenoic acid, 3-[4-(hexopyranosyloxy)-3-hydroxyphenyl]- (9CI) (CA INDEX NAME)

L9 ANSWER 50 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2003:132965 CAPLUS

DOCUMENT NUMBER:

138:163603

TITLE:

Methods for novel sulfur-containing organic nitrate compounds use in the treatment and prevention of human

diseases and conditions

INVENTOR(S):

Garvey, David S.; Letts, L. Gordon

PATENT ASSIGNEE(S):

Nitromed, Inc., USA

SOURCE:

PCT Int. Appl., 43 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PAT | CENT | NO. | | | KIN | D | DATE | | | APPL | ICAT | ION 1 | NO. | | D | ATE | | |
|-----|------|------|-----|-----|------------|-----|------|---------|-----|------|----------|-------|-----|-----|-----|----------|-----|--|
| | | | | | | - | | | | | - | | | | - | - | | |
| WO | 2003 | 0134 | 32 | | A2 | | 2003 | 0220 | | WO 2 | 002- | US24 | 923 | | 2 | 0020 | 807 | |
| WO | 2003 | 0134 | 32 | | A 3 | | 2003 | 1113 | | | | | | | | | | |
| | W: | ΑE, | AG, | AL, | AM, | AT, | AU, | ΑZ, | BA, | BB, | BG, | BR, | BY, | BZ, | CA, | CH, | CN, | |
| | | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EC, | EE, | ES, | FI, | GB, | GD, | GΕ, | GH, | |
| | | GM. | HR. | HU. | ID. | IL. | TN. | TS. | JP. | KE. | KG. | ΚÞ | ΚÞ | K7. | T.C | T.K | T.P | |

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LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
             PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
             UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
             KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
             FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF,
             CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     CA 2453433
                                            CA 2002-2453433
                          A1
                                20030220
                                                                     20020807
                                            EP 2002-786354
     EP 1414432
                                20040506
                                                                     20020807
                          A2
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK
     JP 2005501060
                                20050113
                                             JP 2003-518446
                          т
                                                                     20020807
                          A1
     US 2004152753
                                20040805
                                             US 2004-760672
                                                                     20040121
PRIORITY APPLN. INFO.:
                                             US 2001-311715P
                                                                 P
                                                                    20010810
                                             WO 2002-US24923
                                                                    20020807
```

OTHER SOURCE(S): MARPAT 138:163603

The invention describes methods of use for an organic nitrate compound, or a pharmaceutically acceptable salt thereof, wherein the organic nitrate compound comprises at least one sulfur atom and/or at least one disulfide group. The invention also provides methods for treating, preventing and/or reducing inflammation, pain, and fever; for decreasing or reversing the gastrointestinal, renal and other toxicities resulting from the use of nonsteroidal antiinflammatory compds.; for treating and/or preventing gastrointestinal disorders; for treating inflammatory disease states and disorders; for treating and/or preventing ophthalmic diseases or disorders; for treating and/or improving the gastrointestinal properties of COX-2 inhibitors; for facilitating wound healing; for treating and/or preventing other disorders resulting from elevated levels of cyclooxygenase-2; for decreasing the recurrence of ulcers; for improving gastroprotective properties, anti-Helicobacter pylori properties or antacid properties of proton pump inhibitors; for treating Helicobacter pylori and viral infections. For improving gastroprotective properties of Hz receptor antagonists; for treating and/or preventing inflammations and microbial infections, multiple sclerosis, and viral infections; for treating or preventing restenosis, autoimmune diseases, pathol. conditions resulting from abnormal cell proliferation, polycystic kidney disease, inflammatory diseases or to inhibit wound contraction; for treating or preventing sexual dysfunctions in males and females, for enhancing sexual responses in males and females; for treating or preventing benign prostatic hyperplasia, hypertension, congestive heart failure, variant (Printzmetal) angina , glaucoma, neurodegenerative disorders, vasospastic diseases, cognitive disorders, urge incontinence, and overactive bladder; for reversing the state of anesthesia. For treating or preventing diseases induced by the increased metabolism of cyclic guanosine 3',5'-monophosphate (cGMP); for treating respiratory disorders and for treating neurol. conditions.

IT 53902-12-8, Tranilast

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(methods for novel sulfur-containing organic nitrate compds. use in the treatment and prevention of human diseases and conditions)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

L9 ANSWER 51 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2006:100738 CAPLUS

DOCUMENT NUMBER: 144:198849

TITLE: Novel dosage form comprising modified-release and

immediate-release active ingredients

INVENTOR(S): Vaya, Navin; Karan, Rajesh Singh; Sadanand, Sunil;

Gupta, Vinod Kumar

PATENT ASSIGNEE(S): India

SOURCE: U.S. Pat. Appl. Publ., 49 pp., Cont.-in-part of U.S.

Ser. No. 630,446.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|------|----------|------------------|------------|
| | | | | |
| US 2006024365 | A1 | 20060202 | US 2005-134633 | 20050519 |
| IN 193042 | A1 | 20040626 | IN 2002-MU697 | 20020805 |
| US 2004096499 | A1 | 20040520 | US 2003-630446 | 20030729 |
| PRIORITY APPLN. INFO.: | | | IN 2002-MU697 A | 20020805 |
| | | | IN 2002-MU699 A | 20020805 |
| | | | IN 2003-MU80 A | 20030122 |
| | | | IN 2003-MU82 A | 20030122 |
| | | | US 2003-630446 A | 2 20030729 |

AB A dosage form comprising of a high dose, high solubility active ingredient as modified release and a low dose active ingredient as immediate release where the weight ratio of immediate release active ingredient and modified release active ingredient is from 1:10 to 1:15000 and the weight of modified release active ingredient per unit is from 500 mg to 1500 mg; a process for preparing the dosage form. Tablets containing 10 mg sodium pravastatin and 1000 mg niacin were prepared The release of sodium pravastatin after 24 h was 67.7%, and the release of niacin after 1 h was 84.1%.

IT 5588-21-6, Cintriamide 53902-12-8, Tranilast

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (novel dosage form comprising modified-release and immediate-release
 active ingredients)

RN 5588-21-6 CAPLUS

CN 2-Propenamide, 3-(3,4,5-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

L9 ANSWER 52 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:1290025 CAPLUS

DOCUMENT NUMBER: 144:36329

TITLE: Thiazole compounds as PPAR modulators, their

preparation, pharmaceutical compositions, and use in

therapy

INVENTOR(S): Epple, Robert; Cow, Christopher; Xie, Yongping; Wang,

Xing; Russo, Ross; Azimioara, Mihai; Saez, Enrique

PATENT ASSIGNEE(S): IRM LLC, Bermuda

SOURCE: PCT Int. Appl., 187 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

GΙ

| PAT | rent : | NO. | | | KIN | D | DATE | | | APPL | ICAT | ION I | NO. | | D. | ATE | |
|----------|--------|------|------|-----|-----|-----|------|-------|-----|------|-------|-------|-----|-----|-----|------|-----|
| WO | 2005 | 1160 | 00 | | A1 | _ | 2005 | 1208 | | WO 2 | 005-1 | US18 | 167 | | 2 | 0050 | 524 |
| | W: | ΑE, | AG, | AL, | AM, | AT, | AU, | ΑZ, | BA, | BB, | BG, | BR, | BW, | BY, | ΒZ, | CA, | CH, |
| | | CN, | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EC, | EE, | EG, | ES, | FI, | GB, | GD, |
| | | GE, | GH, | GM, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | KΕ, | KG, | KM, | ΚP, | KR, | ΚZ, |
| • | | LC, | LK, | LR, | LS, | LT, | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | ΜZ, | NA, |
| | | NG, | NI, | NO, | NZ, | OM, | PG, | PH, | ΡL, | PT, | RO, | RU, | SC, | SD, | SE, | SG, | SK, |
| | | SL, | SM, | SY, | ТJ, | TM, | TN, | TR, | TT, | TZ, | UA, | UG, | US, | UZ, | VC, | VN, | YU, |
| | | ZA, | ZM, | zw | | | | | | | | | | | | | |
| | RW: | BW, | GH, | GM, | KE, | LS, | MW, | MZ, | NA, | SD, | SL, | SZ, | TZ, | UG, | ZM, | ZW, | AM, |
| | | ΑZ, | BY, | KG, | KZ, | MD, | RU, | TJ, | TM, | AT, | BE, | BG, | CH, | CY, | CZ, | DΕ, | DK, |
| | | EE, | ES, | FI, | FR, | GB, | GR, | HU, | ΙE, | IS, | IT, | LT, | LU, | MC, | NL, | PL, | PT, |
| | | RO, | SE, | SI, | SK, | TR, | BF, | ВJ, | CF, | CG, | CI, | CM, | GA, | GN, | GQ, | GW, | ML, |
| | | MR, | ΝE, | SN, | TD, | TG | | | | | | | | | | | |
| , AU | 2005 | 2479 | 31 | | A1 | | 2005 | 1208 | | AU 2 | 005-: | 2479: | 31 | | 2 | 0050 | 524 |
| CA | 2563 | 818 | | | A1 | | 2005 | 1208 | 1 | CA 2 | 005-3 | 2563 | 818 | | 2 | 0050 | 524 |
| EP | 1748 | 993 | | | A1 | | 2007 | 0207 | | EP 2 | 005- | 7541 | 30 | | 2 | 0050 | 524 |
| | R: | ΑT, | BE, | ВG, | CH, | CY, | CZ, | DE, | DK, | EE, | ES, | FI, | FR, | GB, | GR, | HU, | ΙE, |
| | | IS, | ΙT, | LI, | LT, | LU, | MC, | NL, | PL, | PT, | RO, | SE, | SI, | SK, | TR | | |
| PRIORITY | APP | LN. | INFO | . : | | | | | • | US 2 | 004- | 5741 | 37P | : | P 2 | 0040 | 524 |
| | | | | | | | | | • | US 2 | 005- | 6489 | 85P | : | P 2 | 0050 | 131 |
| - | | | | | | | | | | WO 2 | 005-1 | US18: | 167 | Ţ | W 2 | 0050 | 524 |
| OTHER SO | OURCE | (S): | | | MAR | PAT | 144: | 36329 | 9 | | | | | | | | |

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

The invention relates to thiazole compds. of formula I, which are modulators of peroxisome proliferator-activated receptors (PPAR), particularly PPARδ. In compds. I, p is 0-3; L is selected from -XOX-, -XS(O)mX-, and -XS(O)mXO-, where m is 0-2 and X is a bond or (un) substituted C1-4 alkylene; R1 is selected from halo, C1-6 alkyl, C1-6 alkoxy, C1-6 hydroxyalkyl, C1-6 haloalkyl, C1-6 haloalkoxy, (un) substituted C6-10 aryl, (un) substituted C5-10 heteroaryl, (un) substituted C3-12 cycloalkyl, and (un) substituted C3-8 heterocyclyl; R2 is -XOXCO2R5 or -XCO2R5, where X is as defined previously and R5 is H or C1-6 alkyl; and R3 and R4 are independently selected from R6 and R6Y, where R6 is (un) substituted C3-12 cycloalkyl, (un) substituted C3-8 heterocyclyl, (un) substituted C6-10 aryl, and (un) substituted C5-13 heteroaryl, and Y is selected from C1-6 alkylene, C2-6 alkenylene, C2-6 alkynylene, -C(0)N(R5)-, and -OX-, where X and R5 are as defined previously, or R3 and R4, together with the atoms to which they are attached, form fused bi- or tricyclic C5-14 heteroaryl; including

pharmaceutically acceptable salts, hydrates, solvates, isomers, and prodrugs thereof. The invention also relates to the preparation of I, pharmaceutical compns. comprising a therapeutically effective amount of compound I in combination with one or more pharmaceutically acceptable excipients, as well as to the use of the compns. to treat or prevent diseases or disorders associated with PPAR activity. Cyclocondensation of 2-bromo-4'-methoxyacetophenone with thioacetamide followed by bromination, demethylation, and alkylation with iso-Pr iodide gave bromothiazole II, which was brominated and substituted with phenol III (preparation in 3 steps from 4-hydroxy-3-methylacetophenone given) to give thiazole IV. Compound IV underwent Suzuki coupling with 4-(trifluoromethoxy)phenylboronic acid and ester hydrolysis to give thiazole V. Most preferred compds. of the invention express an EC50 value for PPARδ of less than 100 nM. compds. of the invention are at least 100-fold selective for PPAR8 over PPARy.

ΤТ 870524-40-6P

> RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(drug candidate; preparation of thiazole compds. as PPAR modulators and their use for treatment and prevention of diseases associated with PPARδ activity)

RN 870524-40-6 CAPLUS

CN 2-Propenoic acid, 3-[3-methoxy-4-[[4-(4-methoxyphenyl)-5-[4-(trifluoromethyl)phenyl]-2-thiazolyl]methoxy]phenyl]- (9CI) (CA INDEX NAME)

MeO
$$\sim$$
 CH $_{\rm CH}$ CH $_{\rm CC}$ CH $_{\rm CH}$ CH $_{\rm CC}$ CH $_{\rm CO}$ CH $_{\rm CH}$ CH $_{\rm CO}$ CH $_{\rm CO$

REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 53 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2000:282025 CAPLUS

DOCUMENT NUMBER:

132:278927

TITLE:

Preparation of N3-(5-methyl-3-isoxazolyl)-3,4-

INVENTOR (S):

dihydroxycinnamide

Ji, Xiaoshen; Wang, Feng; Liu, Min; Jin, Tao; Wang, Jingyuan; Yang, Lianchun; Li, Fei; Lu, Min; Hao, Yong;

Cheng, Yanliang

PATENT ASSIGNEE(S):

SOURCE:

Air Force General Hospital, PLA, Peop. Rep. China Faming Zhuanli Shenqing Gongkai Shuomingshu, 8 pp.

CODEN: CNXXEV

DOCUMENT TYPE:

Patent Chinese

LANGUAGE:

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE -----------19981230 CN 1203230 Α CN 1998-101434 19980428 CN 1100047 В 20030129

PRIORITY APPLN. INFO.:

19980428

OTHER SOURCE(S):

CASREACT 132:278927

The title compound is prepared by acylating 3-amino-5-methyl-isoxazole with 3,4-dihydroxycinnamic acid in anhydrous THF and in the presence of DCCI overnight, filtering, and separating with vacuum column and EtOAc-acetone as gradient eluent. The title compound may be prepared by chlorinating 3,4-dihydroxycinnamic acid with SOCl2 in solvent and/or in the presence of HMPA for 3-4 h, and acylating 3-amino-5-methyl-isoxazole for 4 h. The title compound may be prepared by allowing to react 3,4- dihydroxycinnamic acid with POCl3 and 4-nitrophenol in benzene for 2 h, and substituting 3-amino-5-methyl-isoxazole for 2 h. The tablet as endothelial element antagonist is composed of the compound 40, starch 20, alginic acid 20, Na alginate 20, and Mg stearate 1.3 mg.

IT 227275-60-7P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of N3-(5-methyl-3-isoxazolyl)-3,4-dihydroxycinnamide as endothelin 1 antagonist)

RN 227275-60-7 CAPLUS

CN 2-Propenamide, 3-(3,4-dihydroxyphenyl)-N-(5-methyl-3-isoxazolyl)- (9CI) (CA INDEX NAME)

IT 331-39-5, 3,4-Dihydroxycinnamic acid

RL: RCT (Reactant); RACT (Reactant or reagent) (preparation of N3-(5-methyl-3-isoxazolyl)-3,4-dihydroxycinnamide as endothelin 1 antagonist)

RN 331-39-5 CAPLUS

CN 2-Propenoic acid, 3-(3,4-dihydroxyphenyl)- (9CI) (CA INDEX NAME)

L9 ANSWER 54 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1995:888203 CAPLUS

DOCUMENT NUMBER:

124:75939

TITLE:

Inhibition of PDGF- and TGF-β1-induced collagen

synthesis, migration and proliferation by tranilast in

vascular smooth muscle cells from spontaneously

hypertensive rats

AUTHOR (S):

Miyazawa, Keiji; Kikuchi, Shinji; Fukuyama, Juichi;

Hamano, Shuichiro; Ujiie, Arao

CORPORATE SOURCE:

Pharmacological Laboratories, Kissei Pharmaceutical

Co. Ltd., Hotaka, Nagano, 399-83, Japan

SOURCE:

Atherosclerosis (Shannon, Ireland) (1995), 118(2),

213 - 21

CODEN: ATHSBL; ISSN: 0021-9150

PUBLISHER: Elsevier DOCUMENT TYPE: Journal English LANGUAGE:

Vascular smooth muscle cells (VSMC) from spontaneously hypertensive rats (SHR) proliferate faster and are more sensitive to transforming growth factor- β 1 (TGF- β 1) than those of normotensive Wistar-Kyoto rats. We studied the in vitro effects of tranilast, an anti-allergic drug, on the proliferation, migration and extracellular matrix synthesis in the SHR-VSMC. There were many inhibitory effects of tranilast (30-300 μM) on SHR-VSMC. One is the effect on the proliferation stimulated with fetal bovine serum (FBS), TGF- β 1 and platelet-derived growth factor-BB (PDGF-BB). Another is the effect on the PDGF-BB-induced migration. Lastly, tranilast exhibited

inhibitory effects on spontaneous collagen synthesis and

TGF-β1-induced collagen and glycosaminoglycan synthesis. Collagen induced the VSMC migration concentration-dependently. These results suggest

that

tranilast may prevent restenosis after percutaneous transluminal coronary angioplasty.

TT 53902-12-8, Tranilast

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES

(inhibition of vascular smooth muscle proliferation by tranilast and possible prevention of restenosis after angioplasty)

RN 53902-12-8 CAPLUS

Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) CN (CA INDEX NAME)

ANSWER 55 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1993:468088 CAPLUS

DOCUMENT NUMBER: 119:68088

TITLE: Caracasanamide, a novel hypotensive agent from

Verbesina caracasana

AUTHOR (S): Delle Monache, Giuliano; Botta, Bruno; Delle Monache,

Franco; Espinal, Romulo; De Bonnevaux, Stella C.; De Luca, Carlo; Botta, Maurizio; Corelli, Federico;

Carmignani, Marco

CORPORATE SOURCE: Cent. Chim. Recett., Rome, 00168, Italy

SOURCE: Bioorganic & Medicinal Chemistry Letters (1992), 2(5),

415-18

CODEN: BMCLE8; ISSN: 0960-894X

DOCUMENT TYPE: Journal LANGUAGE: English

AB The hypotensive agent from Verbesina caracasana is shown to be a novel guanidino-amide which occurs in (Z)- and (E)-forms. The structure of the compound was confirmed by the synthesis of the (E) form.

TΤ 146269-39-8 146269-40-1, (Z)-Caracasanamide

RL: BIOL (Biological study)

(from Verbesina caracasana, structure and hypotensive activity of)

RN 146269-39-8 CAPLUS CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[[imino[(3-methyl-2-butenyl)amino]methyl]amino]butyl]-, (E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 146269-40-1 CAPLUS

CN 2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[4-[[imino[(3-methyl-2-butenyl)amino]methyl]amino]butyl]-, (2Z)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

$$\begin{array}{c|c} & H & H & H \\ \hline Z & O & NH & NH \\ \hline \\ OMe & \\ \end{array}$$

L9 ANSWER 56 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1990:235330 CAPLUS

DOCUMENT NUMBER: 112:235330

TITLE: Preparation of 3,4-dihydro-2-[(2-substituted)phenyl]-

2H-1,4-benzothiazin-3-ones as calcium antagonists

APPLICATION NO.

DATE

PATENT ASSIGNEE(S): Hoechst A.-G., Fed. Rep. Ger.

KIND

SOURCE: Austrian, 22 pp.

CODEN: AUXXAK

DATE

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.

| AT 389112 | В | 19891025 | AT 1987-2585 | 19871008 |
|-----------------|-----------------|--------------|--------------------------------|-----------------|
| AT 8702585 | A | 19890315 | | |
| PRIORITY APPLN. | INFO.: | | AT 1987-2585 | 19871008 |
| OTHER SOURCE(S) | : CASRE | ACT 112:2353 | 30; MARPAT 112:235330 |) |
| GI For diagram | m(s), see print | ed CA Issue. | | |
| AB The title | compds. [I; R1, | R6, R7 = H, | C1-4 alkyl, C1-3 alk | oxy, F, Cl, Br, |
| CF3, NO2, 0 | OH, AcNH, amino | ; R2 = H, C1 | -10 alkyl, C3-10 alke | envl. |
| | | | 15 alkyl, C3-15 alker | |
| | | | initions of R1 except | |
| | | |)n; $X2 = NR9$, $Q1$, $Q2$, | |
| | | | (CH2)q, CH:CH, etc.; | |
| | | | ; R9 = H, C1-10 alkyl | |
| | | | m = 1-4; $n = 2$, 3; p | |
| | | | | |
| | | | ally acceptable salts | |
| | | | ment of angina pector | |
| arrhythmia | , and hypertens | ion. A mixt | ure of 3,4-dihydro-2- | |
| | | | | |

isopropyl-4-methyl-2-[2-(4-bromobutoxy)phenyl]-2H-1,4-benzothiazin-3-one 6.68, K2CO3 2.07, and 3,4,5-trimethoxyphenylacetic acid piperazide (preparation given) 6.63 g was refluxed 5 h in DMF to give 2.4 g title compound II which was converted to 1.7 g of II.HCl. I in vitro antagonized Ca with IC50 values of 10-6 - 10-10 M.

IT 90-50-6, 3,4,5-Trimethoxycinnamic acid

RL: RCT (Reactant); RACT (Reactant or reagent)

(reaction of, in preparation of calcium antagonist)

RN 90-50-6 CAPLUS

CN 2-Propenoic acid, 3-(3,4,5-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

L9 ANSWER 57 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1976:44513 CAPLUS

DOCUMENT NUMBER: 84:44513

TITLE: Protoberberine derivatives

INVENTOR(S):
Kametani, Tetsuji

PATENT ASSIGNEE(S): Japan Chemipha Co., Ltd., Japan SOURCE: Jpn. Kokai Tokkyo Koho, 4 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|------|----------|-----------------|----------|
| | | | | - |
| JP 50101399 | Α | 19750811 | JP 1974-8847 | 19740119 |
| PRIORITY APPLN. INFO.: | | | JP 1974-8847 | 19740119 |

GI For diagram(s), see printed CA Issue.

AB Protoberberine derivs. (I; R1 = lower alkyl; R2 = straight chain or branched lower alkyl, pyridyl, phenyl or phenylvinyl substituted with 2 or 3 lower alkoxy groups) were prepared by reaction of protoberberines II with carboxylic acids R2CO2H or their reactive derivs. I had analgesic, vasodilating, and hypotensive activities (no data). Thus, Ac2O was added to 3.8 g II (R1 = Me) to give 51.3% I (R1 = R2 = Me). Also were prepared I (R1, R2 given): Me, tert-Bu; Me, 3-pyridyl; Me, 3,4,5-(MeO)3C6H2; Me, 3,4-(MeO)2C6H3CH:CH; and Me, 3,4,5-(MeO)3C6H2CH:CH.

IT 90-50-6 2316-26-9

RL: RCT (Reactant); RACT (Reactant or reagent)
 (acylation by, of hydroxyprotoberberines)

RN 90-50-6 CAPLUS

CN 2-Propenoic acid, 3-(3,4,5-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

RN 2316-26-9 CAPLUS

CN 2-Propenoic acid, 3-(3,4-dimethoxyphenyl)- (9CI) (CA INDEX NAME)

L9 ANSWER 58 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1975:458672 CAPLUS

DOCUMENT NUMBER: 83:58672

TITLE: 4-Biphenylyl isoquinoline derivatives

INVENTOR(S): Jansen, Alexander Bertus A.; Hollywood, John; Wilson,

Alan Brian

PATENT ASSIGNEE(S): UK

SOURCE: U.S., 6 pp.

CODEN: USXXAM DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|------------------------|------|----------|-----------------|---|----------|
| | | | | | |
| US 3823148 | A | 19740709 | US 1972-256955 | | 19720525 |
| GB 1386076 | A | 19750305 | GB 1971-18765 | | 19720602 |
| PRIORITY APPLN. INFO.: | | | GB 1971-18765 | A | 19710603 |

GI For diagram(s), see printed CA Issue.

The dihydroisoquinoline I (R = p-PhC6H4, 1-adamantyl, p-MeSO2NHC6H4CH2, p-H2NC6H4CH2, etc.; R1 = H, Me) were prepared by cyclization of amides. Thus, p-PhC6H4COCl was treated with 3,4-(MeO)2C6H3CH2CH2NH2 to give 3,4-(MeO)2C6H3CH2CH2NHCOC6H4Cl-p, which was cyclized with POCl3 to give I (R = p-PhCl6H4, R1 = Me). Several I were reduced to the 1,2,3,4-tetrahydro derivs. I were hypotensives, depressants, and anticonvulsants (no data).

IT 56205-55-1P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation and cyclization of)

RN 56205-55-1 CAPLUS

$$\begin{array}{c|c} \text{OMe} \\ \text{O$$

IT 90-50-6

RL: RCT (Reactant); RACT (Reactant or reagent)
 (reaction of, with amines)

RN 90-50-6 CAPLUS

CN 2-Propenoic acid, 3-(3,4,5-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

L9 ANSWER 59 OF 98. CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1968:402921 CAPLUS

DOCUMENT NUMBER: 69:2921

TITLE: Synthesis and pharmacology of a series of amide

derivatives of 3,4,5-trimethoxycinnamic acid and their

analog

AUTHOR(S): Cerbai, Guido; Turbanti, L.; Bianchini, P.; Bramanti,

Giancarlo; Tellini, N.

CORPORATE SOURCE: Direzione Ric., Lab. Guidotti and C. S.p.A., Pisa,

Italv

SOURCE: Bollettino Chimico Farmaceutico (1967), 106, 837-54

CODEN: BCFAAI; ISSN: 0006-6648

DOCUMENT TYPE: Journal LANGUAGE: Italian

GI For diagram(s), see printed CA Issue.

The sedative and hypotensive activity of N-(3,4,5-trimethoxycinnamoyl)morpholine (I) (R1 = R2 = MeO, R3 = morpholino) led to the synthesis of a
series of new amide derivs., with changes in both the acid and the basic
part of the mol. Thus, to a solution of 3.6 g. acetylsinapyl chloride and
1.51 g. Et3N in 60 ml. C6H6, 1.13 g. morpholine was added slowly, with
stirring and cooling; stirring was continued 1 hr. at room temperature, and the
mixture refluxed 3-4 hrs. and worked up to give 60% I (R1 = AcO, R2 = MeO,
R3 = morpholino). In preparation of derivs. of acetylferulic acid, oily
residues were obtained which had to be washed with petroleum ether and
dried in vacuo prior to crystallization. In a somewhat different way, I (R1 =

R2 =

MeO, R3 = 4-oxopiperidino) (Ia) was prepared by decarboxylating 7.5 g. 3-carbethoxy-4-piperidone-HCl in 25 ml. 6N HCl by refluxing the mixture 2 hrs. and evaporating the solvent; the residue was taken up with 5 ml. H2O, 5 g. anhydrous K2CO3, 20 ml. CHCl3, and 9.35 g. 3,4,5-trimethoxycinnamoyl chloride in 50 ml. CHCl3 were added successively, and the mixture stirred 2 hrs., refluxed 4 hrs. and, after addition of 3 ml. EtOH, refluxed again 2 hrs. and worked up to give Ia, m. 135-6° (EtOH); p-nitrophenylhydrazone m. 195-7°. I prepared are given in the table. I (R1 = AcO, R2 = MeO, R3 = morpholino) (2.1 g.) was dissolved in 25 ml. absolute MeOH, to the cooled solution MeONa in MeOH (prepared from 0.15 g. Na in 6 ml. MeOH) was added slowly with stirring, and the mixture kept overnight at room temperature and worked up to give .apprx.40% the following I (R1 = OH) (R2, R3, and m.p. given): H, morpholino, 172-4°; H, 2-methylmorpholino, 89-91°; H, 2-ethylmorpholino, 64-6°; H, 3-ethylmorpholino, 79-81°; MeO, morpholino, 201-2°; and MeO, 2-methylmorpholino, 155°. In a similar way the following 3,4,5-trimethoxyacetic acid derivs. (II) were prepared (R given): morpholino (b0.1 207-9°); and pyrrolidino (m. 64-5°). A solution of 7.48 g. ethylene oxide in 25 ml. EtOH was added to a solution of 20 g. BuCH(NH2)CH2OH in 15 ml. EtOH with stirring and cooling to 0°, stirring continued 2 hrs., and the mixture kept 24 hrs. at room temperature, heated 1 hr. on a water bath, and worked up to give 17.8 g. N-(2-hydroxyethyl)-2-aminohexanol (III), b9.1 114-15°. In the same way PrCH(NH2)CH2OH gave N-(2-hydroxyethyl)-2-aminopentanol (IV), b0.7-0.8 119-22°. III (10 g.) was added dropwise with stirring to 7 ml. concentrated H2SO4 at 0°, and the mixture heated 6 hrs. at 150-60° (distillation of H2O formed) and worked up to give 3.9 g. 3-butylmorpholine

b. 198°; HCl salt m. 133° (absolute EtOH). V and VI were used

in the preparation of the corresponding I. The results of the pharmacological screening made with the described I can be summarized as follows: (a) the introduction at C-2 of the morpholine ring of a Me and Et group increased the sedative and decreased the antihistaminic activity of I; (b) the introduction of Et, Br, and Bu at C-3 caused appearance of hypotensive action; (c) the introduction of Me at C-2 and C-6 induced changes in the antiedemic activity and potentiated the action on the bronchospasm induced by histamine; (d) the elimination of the MeO-groups and acetylation in C-4 of the C6H6-ring increased the analgesic action of I. 5588-21-6DP, Cinnamamide, 3,4,5-trimethoxy-, derivs.

TΤ 16562-68-8P

RL: SPN (Synthetic preparation); PREP (Preparation)

(preparation and pharmacology of)

5588-21-6 CAPLUS RN

2-Propenamide, 3-(3,4,5-trimethoxyphenyl)- (9CI) (CA INDEX NAME) CN

RN 16562-68-8 CAPLUS

2-Propenamide, N-(2-hydroxyethyl)-N-(2-hydroxypropyl)-3-(3,4,5-CN trimethoxyphenyl) - (9CI) (CA INDEX NAME)

L9 ANSWER 60 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2005:49212 CAPLUS

DOCUMENT NUMBER:

142:367143

TITLE:

Predictive factors for ischemic target vessel

revascularization in the Prevention of Restenosis with

Tranilast and its Outcomes (PRESTO) trial

AUTHOR (S):

Singh, Mandeep; Gersh, Bernard J.; McClelland, Robyn

L.; Ho, Kalon K. L.; Willerson, James T.; Penny,

William F.; Holmes, David R.

CORPORATE SOURCE:

Division of Internal Medicine and Cardiovascular

Diseases, Mayo Clinic and Mayo Foundation, Rochester,

MN, USA

SOURCE:

Journal of the American College of Cardiology (2005),

45(2), 198-203

CODEN: JACCDI; ISSN: 0735-1097

PUBLISHER:

Elsevier Inc.

DOCUMENT TYPE:

Journal LANGUAGE: English

The aim of the present study was to determine the rates of target vessel revascularization (TVR) and to determine predictors of TVR from clin. and angiog. variables available in the Prevention of Restenosis With Tranilast and its Outcomes (PRESTO) database. The rates of TVR after percutaneous revascularization procedures, and its prediction with available clin. and

angiog. variables, is less well known. We studied nine-month TVR in 11,484 patients enrolled in the PRESTO trial. Clin., lesion-related, and procedural characteristics were analyzed in a logistic regression model. Study data were divided at random into an 80% training set on which the models were developed and a 20% hold-out set on which the model properties were evaluated. A total of 14% (n = 1,609) had ischemic TVR. Clin. variables with increased risk for TVR included younger age; hypertension; diabetes mellitus; nonsmokers; unstable angina; previous coronary artery bypass grafting; peripheral vascular disease; procedure- and lesion-related such as ostial location, multilesion angioplasty, location in the left anterior descending artery, length ≥20 mm, in-stent restenosis at baseline, and use of rotablator. There was significant increase in the risk of ischemic TVR at U.S. treatment sites. Smoking and stent placement were associated with lower risk of ischemic TVR. The mean area (± SD) under the receiver-operating characteristic curve of the bootstrap samples was 0.66, indicating a modest ability of the model to discriminate patients who needed TVR on follow-up. Despite being the largest prospective trial designed to test restenosis, the discriminatory ability of the clin. and angiog. variables to predict TVR is modest.

IT 53902-12-8, Tranilast

> RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(PRESTO trial showed modest accuracy of clin. and angiog. variables for prediction of target vessel revascularization in percutaneous coronary intervention patient)

RΝ 53902-12-8 CAPLUS

Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) CN (CA INDEX NAME)

REFERENCE COUNT: 20 THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 61 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1966:38793 CAPLUS

DOCUMENT NUMBER: 64:38793 ORIGINAL REFERENCE NO.: 64:7248a-b

TITLE: Recent advances in hypotensive pharmaceuticals; dopa

decarboxylase inhibitors Wermuth, Camille Georges

AUTHOR(S): CORPORATE SOURCE: Fac. Pharm., Strasbourg, Fr. SOURCE: Therapie (1965), 20(6), 1569-78

CODEN: THERAP; ISSN: 0040-5957

DOCUMENT TYPE: Journal

LANGUAGE: French

AB Previous studies are reviewed concerning dopa decarboxylase and the use of enzyme inhibitors to reduce hypertension. Intravenous α -methyl dopa, N-(DL-seryl)-N'-(2,3,4-trihydroxybenzyl)hydrazine, piperidine caffeate, and protocatechuic aldoxime inhibited the pressor response induced by 5 mg. DL-dopa in urethanized and hexamethoniumpretreated rats; α -(aminooxy)-6-bromo-m-cresol was not inhibitory. 21 references.

300-51-6, Cinnamic acid, 3,4-dihydroxy-, compound with piperidine IT (1:1)

```
(blood pressure lowering by)
RN
     300-51-6 CAPLUS
     2-Propenoic acid, 3-(3,4-dihydroxyphenyl)-, compd. with piperidine (1:1)
CN
     (9CI) (CA INDEX NAME)
     CM
           1
     CRN
          331-39-5
     CMF
          C9 H8 O4
            CH = CH - CO_2H
HO
      OH
          2
     CM
     CRN
          110-89-4
     CMF
          C5 H11 N
     NH
IT
     300-51-6P, Cinnamic acid, 3,4-dihydroxy-, compound with piperidine
     (1:1)
     RL: PREP (Preparation)
        (preparation of)
RN
     300-51-6 CAPLUS
CN
     2-Propenoic acid, 3-(3,4-dihydroxyphenyl)-, compd. with piperidine (1:1)
     (9CI) (CA INDEX NAME)
     CM
          1
     CRN
          331-39-5
     CMF
          C9 H8 O4
            CH = CH - CO_2H
HO
      OH
     CM
          2
     CRN 110-89-4
CMF C5 H11 N
```



L9 ANSWER 62 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1958:35475 CAPLUS

DOCUMENT NUMBER: 52:35475
ORIGINAL REFERENCE NO.: 52:6405e-g

TITLE: Dimethylaminoethyl esters of polyalkoxy benzoic and

cinnamic acids

DOCUMENT TYPE: Patent

LANGUAGE: Unavailable

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE
US 2816133 19571210 US

3,4,5-(MeO)3C6H2CO2H (7 g.) and 11.9 g. SOCl2 (I) refluxed 2 hrs., the AB excess I evaporated in vacuo, the solid acid chloride (II) dissolved in 15 ml. C6H6 and the C6H6 evaporated in vacuo, the II in 50 ml. PhMe added dropwise with stirring and cooling to 8.9 g. Me2NCH2CH2OH (III) in 25 ml. dry PhMe, the mixture warmed to 100° and kept at 100° an hr., the III.HCl filtered off, the filtrate washed with H2O and dried, the solvent removed, and the residue distilled in vacuo gave 82% 3,4,5-(MeO)3C6H2CO2CH2CH2NMe2, b0.5 155°; HCl salt, m. 126-7° (ether). Similarly prepared were III esters of the following acids: 2,3,4-(MeO)3C6H2CO2H, b0.5 142-6°, n2OD 1.5195 (HCl salt, m. 132-3°); 2,4,6-(MeO)3C6H2CO2H, m. 61-2° (HCl salt, m. 190-°1); 2,3,4-(MeO)3C6H2CH:CHCO2H HCl salt, m. 121-2°; 3,4,5-(MeO)3 analog HCl salt, m. 182-3° (decomposition). These compds. have desirable pharmacol. properties which affect the cardiovascular dynamics of animals. They are nontoxic and useful in treatment of cardiovascular abnormalities such as hypertension. 90-50-6, Cinnamic acid, 3,4,5-trimethoxy-, 2-dimethylaminoethyl

RN 90-50-6 CAPLUS

CN 2-Propenoic acid, 3-(3,4,5-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

RN 33130-03-9 CAPLUS

CN 2-Propenoic acid, 3-(2,3,4-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

L9 ANSWER 63 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2006:950067 CAPLUS

DOCUMENT NUMBER: 145:342435

TITLE: Manufacture and application of sodium ferulate

injection
INVENTOR(S): Guo, Zhihua

PATENT ASSIGNEE(S): Lokis Pharmaceutical (Jilin) Co., Ltd., Peop. Rep.

China

SOURCE: Faming Zhuanli Shenqing Gongkai Shuomingshu, 8pp.

CODEN: CNXXEV

DOCUMENT TYPE: Patent LANGUAGE: Chinese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|----------|-------------|---------------------------|--------------|
| | | | | - |
| CN 1830428 | A | 20060913 | CN 2005-10051479 | 20050308 |
| PRIORITY APPLN. INFO.: | • | | CN 2005-10051479 | 20050308 |
| | | | contains sodium ferulate, | |
| | | | injection water. The in | |
| manufactured by gr | inding t | the mixture | of sodium ferulate and 1 | -40 times |
| propylene | | | | |

glycol homogeneously, dissolving in injection water, adding sodium hydrogen sulfite, ultrafiltering, bottling, and sterilizing to obtain the final product. Sodium ferulate can be prevented from hydrolyzation by propylene glycol and oxidative degradation by sodium hydrogen sulfite. The injection can be used to treat arteriosclerosis, coronary heart disease, cerebrovascular diseases, glomerulus diseases, pulmonary hypertension, vascular diseases resulted from diabetes mellitus and angiitis, leukopenia, thrombocytopenia, migraine, and vascular

IT 24276-84-4, Sodium ferulate

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(manufacture and application of sodium ferulate injection)

RN 24276-84-4 CAPLUS

headache.

CN 2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)-, monosodium salt (9CI) (CA INDEX NAME)

Na

L9 ANSWER 64 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2006:946650 CAPLUS

DOCUMENT NUMBER: 145:342430

TITLE: Method for manufacturing freeze-dried injection

containing sodium ferulate

INVENTOR(S): Guo, Zhihua

PATENT ASSIGNEE(S): Barrymore Pharmaceutical (Tonghua) Co., Ltd., Peop.

Rep. China

SOURCE: Faming Zhuanli Shenqing Gongkai Shuomingshu, 12pp.

CODEN: CNXXEV

DOCUMENT TYPE: LANGUAGE:

ferulate

Patent Chinese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE _____ ------------------------CN 2005-10051477 20060913 Α CN 1830427 20050308 CN 2005-10051477 PRIORITY APPLN. INFO.: 20050308 The title method comprises: (1) dissolving sodium ferulate in water for injection in nitrogen protection and under protecting from light, and ultrafiltering to obtain an ultrafiltrate containing sodium ferulate, (2) dissolving mannitol in water for injection, adding activated carbon, boiling, and filtering for removing activated carbon to obtain a filtrate containing mannitol, and (3) adding the ultrafiltrate containing sodium

to the filtrate containing mannitol, mixing to obtain a uniform solution, fixing

volume, adjusting pH, inspecting quality of semifinished product, sterilizing, packaging in penicillin bottle in nitrogen protection and under protecting from light, freeze-drying, and sealing to obtain the freeze-dried injection containing sodium ferulate. This injection has the advantages of low degradation of sodium ferulate, good solubility and stability.

low stimulation, and stable curative effect. The injection can be used for treating atherosclerosis, coronary heart disease, cerebrovascular disease, glomerular disease, pulmonary hypertension, diabetic angiopathy, angitis, leukopenia, thrombocytopenia, migraine and vascular headache.

IT 24276-84-4, Sodium ferulate

RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); PKT (Pharmacokinetics); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(method for manufacturing freeze-dried injection containing sodium ferulate)

RN 24276-84-4 CAPLUS

CN 2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)-, monosodium salt (9CI) (CA INDEX NAME)

Na

L9 ANSWER 65 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2004:295980 CAPLUS

DOCUMENT NUMBER:

141:325356

TITLE:

Inhibitory influences of xanthine oxidase inhibitor and angiotensin I-converting enzyme inhibitor on multinucleated giant cell formation from monocytes by down-regulation of adhesion molecules and purinergic

receptors

AUTHOR(S):

Mizuno, K.; Okamoto, H.; Horio, T.

CORPORATE SOURCE:

Department of Dermatology, Kansai Medical University,

Moriguchi, Osaka, 570-8507, Japan

SOURCE: British Journal of Dermatology (2004), 150(2), 205-210

CODEN: BJDEAZ; ISSN: 0007-0963

PUBLISHER: Blackwell Publishing Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

Allopurinol, a xanthine oxidase inhibitor, and captopril, an inhibitor of angiotensin I-converting enzyme, are widely used for hyperuricemia and hypertension, resp. There have been reported cases showing that these two agents are effective for the treatment of granulomatous diseases such as sarcoidosis, although the mode of action is not elucidated. We examined the in vitro effects of these agents on the formation of multinucleated giant cells (MGC) from human monocytes by Con A-stimulated mononuclear cell supernatants (conditioned medium). We cultured monocytes with conditioned medium and each agent and compared the rate of MGC formation as well as the expression of adhesion mols. and P2X7 receptor, which are involved in MGC formation. The addition of 25 or 100 µg mL-1 allopurinol or 0.125-1.0 µg mL-1 captopril inhibited MGC formation. Monocytes treated with these agents exhibited less expression of intercellular adhesion mol.-1 (ICAM-1) than untreated monocytes. susceptibility of monocytes cultured in conditioned medium for 24 h to 2'-and 3'-o-(4-benzoyl-benzoyl)ATP-mediated cytolysis was significantly lower in monocytes treated with these agents than in untreated monocytes. Allopurinol and captopril have a therapeutic effect on granulomatous disorders by a direct action on monocyte/macrophage lineage cells partly through down-regulation of ICAM-1 and P2X7 receptor.

IT 53902-12-8, Tranilast

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(tranilast inhibited MGC formation in dose dependent manner, but effect was less than allopurinol or captopril and did not significantly affect LDH release on human monocyte)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

REFERENCE COUNT: 28 THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 66 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2002:694488 CAPLUS

DOCUMENT NUMBER: 138:82828

AUTHOR (S):

TITLE: Left ventricular hypertrophy: a new approach for

fibrosis inhibition Muiesan, Maria Lorenza

CORPORATE SOURCE: Dipartimento di Scienze Mediche, Medicina, Spedali

Civili, Brescia, 25100, Italy

SOURCE: Journal of Hypertension (2002), 20(4), 611-613

CODEN: JOHYD3; ISSN: 0263-6352 Lippincott Williams & Wilkins

PUBLISHER: Lippincott Williams & Wi DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

AB A review on the effect of treatment with translast for 12 wk in rats with renovascular hypertension compared to an angiotensin-converting

enzyme (ACE)-inhibitor or no treatment. Tranilast was found to reverse

transforming growth factor- β 1 mediated cardiac fibrosis, independently of blood pressure, while treatment with an ACE-inhibitor reduced blood pressure and left ventricular fibrosis. This result provides evidence that TGF- β 1 plays a major role in the process of cardiac fibrosis, particularly when the renin-angiotensin system is activated.

IT 53902-12-8, Tranilast

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(tranilast for treatment of left ventricular hypertrophy)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

REFERENCE COUNT: 28 THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 67 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2006:889492 CAPLUS

DOCUMENT NUMBER:

145:299764

TITLE:

Implantable small percutaneous valve and methods of

delivery

INVENTOR(S):

Kheradvar, Arash; Ravichandran, Guruswami; Gharib,

Morteza

PATENT ASSIGNEE(S):

USA

SOURCE:

U.S. Pat. Appl. Publ., 24pp.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | KII | | | APPL | CATION 1 | DATE | | |
|-------------------|-------------|---------|-------|---------|----------|--------|---------------|--|
| US 200619518 | | | 0831 | US 20 | 006-3618 | 50 | 20060224 | |
| WO 200611022 | 28 A: | 2006 | 1019 | WO 20 | 006-US70 | 22 | 20060227 | |
| W: AE, | AG, AL, AM | AT, AU, | AZ, E | BA, BB, | BG, BR, | BW, BY | , BZ, CA, CH, | |
| | | | | | | | , FI, GB, GD, | |
| | | | | | | | , KN, KP, KR, | |
| KZ, | LC, LK, LR | LS, LT, | LU, I | LV, LY, | MA, MD, | MG, MK | , MN, MW, MX, | |
| | | | | | | | , SC, SD, SE, | |
| SG, | SK, SL, SM | SY, TJ, | TM, T | TN, TR, | TT, TZ, | UA, UG | , US, UZ, VC, | |
| VN, | YU, ZA, ZM | ZW | | | | | | |
| RW: AT, | BE, BG, CH | CY, CZ, | DE, I | DK, EE, | ES, FI, | FR, GB | , GR, HU, IE, | |
| IS, | IT, LT, LU | LV, MC, | ŇL, F | PL, PT, | RO, SE, | SI, SK | , TR, BF, BJ, | |
| CF, | CG, CI, CM, | GA, GN, | GQ, G | GW, ML, | MR, NE, | SN, TD | , TG, BW, GH, | |
| GM, | KE, LS, MW, | MZ, NA, | SD, S | SL, SZ, | TZ, UG, | ZM, ZW | , AM, AZ, BY, | |
| KG, | KZ, MD, RU, | TJ, TM | | | | | | |
| PRIORITY APPLN. 1 | NFO.: | | | US 20 | 05-6564 | 56P | P 20050225 | |
| | | | | US 20 | 05-6574 | 74P | P 20050301 | |
| | | | | US 20 | 05-74834 | 15P | P 20051206 | |
| | | | | US 20 | 06-75670 | 05P | P 20060106 | |
| 7D 7 | | | | US 20 | 06-3618 | 50 | A 20060224 | |

AB An implantable prosthetic valve system for implantation in a body channel,

more particularly, to an implantable prosthetic heart valve suitable for replacement of a defect or diseased human heart valve and method of delivery are provided. The prosthetic valve is transformable from a first helical pre-implantation configuration to a second valvular functional configuration. The valve comprises a support structure with leaflets made from synthetic material, engineered biol. tissue, biol. valvular leaflet tissue, pericardial tissue, and crosslinked pericardial tissue. At least a portion of the support structure and the leaflets is covered with cloth. The support structure comprises a circular stent made, e.g., of shape memory Nitinol and a plurality of elongate support arms. The prosthetic valve system comprises a radially or helically collapsible and expandable crown loaded with at least one bioactive agent.

IT 53902-12-8, Tranilast

RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(implantable small percutaneous valve and delivery apparatus)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

L9 ANSWER 68 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2006:666025 CAPLUS

DOCUMENT NUMBER: 145:152690

TITLE: Method for inducing crystalline state transition in

pharmaceuticals

INVENTOR(S): Nakamichi, Kouichi; Izumi, Shougo; Oka, Masaaki

PATENT ASSIGNEE(S): Nippon Shinyaju Company, Ltd., Japan

SOURCE: U.S., 18 pp., Cont.-in-part of U.S. 5,456,923.

CODEN: USXXAM

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

| PATENT NO | • | KIND | DATE | APPLICATION NO. | DATE |
|----------------|------------|------------|-----------|---------------------|--------------------|
| US 581154 | 7 | А | 19980922 | US 1995-416815 | 19950609 |
| CA 214727 | 9 | A1 | | CA 1993-2147279 | |
| WO 940856 | 1 | A1 | 19940428 | WO 1993-JP1469 | 19931013 |
| W: A | U, BR, CA, | FI, HU | , JP, KR, | NO, NZ, RU, US | |
| RW: A | T, BE, CH, | DE, DK | , ES, FR, | GB, GR, IE, IT, LU, | MC, NL, PT, SE |
| AU 935160 | 7 | A | 19940509 | AU 1993-51607 | 19931013 |
| EP 665009 | | A1 | 19950802 | EP 1993-922625 | 19931013 |
| EP 665009 | | B1 | 20000216 | | |
| R: A | T, BE, CH, | DE, DK | , ES, FR, | GB, GR, IE, IT, LI, | LU, MC, NL, PT, SE |
| AT 189770 | | T | 20000315 | AT 1993-922625 | 19931013 |
| ES 214506 | 3 | T 3 | 20000701 | ES 1993-922625 | 19931013 |
| US 545692 | 3 | A | 19951010 | US 1993-129133 | 19931115 |
| PRIORITY APPLN | . INFO.: | | | JP 1992-303085 | A 19921014 |
| | | | | WO 1993-JP1469 | W 19931013 |
| | | | | US 1993-129133 | A2 19931115 |
| | | | | JP 1991-112554 | |
| | | | | WO 1992-JP470 | W 19920414 |

This invention has for its object to provide a method of inducing a AB transition in crystalline state of a crystallizable pharmaceutical with great ease and improved efficiency and uniformity on a high production scale. An extruder is used for inducing a transition from one crystalline state (A) to another crystalline state in a crystallizable pharmaceutical. An extruded indomethacin (form α) was converted to an amorphous form.

IT 53902-12-8, Tranilast

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES

(method for inducing crystalline state transition in pharmaceuticals)

RN53902-12-8 CAPLUS

Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) CN (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS 10 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

CAPLUS COPYRIGHT 2007 ACS on STN L9ANSWER 69 OF 98

ACCESSION NUMBER:

2006:151208 CAPLUS

DOCUMENT NUMBER:

144:219324

TITLE:

Transnasal composition having immediate action and

high absorbability

INVENTOR(S):

Nagata, Ryoichi; Haruta, Shunji

PATENT ASSIGNEE(S):

Translational Research, Ltd., Japan

SOURCE:

PCT Int. Appl., 29 pp.

DOCUMENT TYPE:

CODEN: PIXXD2

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT | NO. | | | KIN | D : | DATE | | | APPL | ICAT | ION I | NO. | | D | ATE | |
|---------|--------------|-----|-----|-----|-----|------|--------------|-----|------|------|-------|-----|-----|-----|------|-----|
| | - | | | | _ | | - | | | | | | | - | | |
| WO 2006 | 0165 | 30 | | A1 | | 2006 | 0216 | 1 | WO 2 | 005- | JP14: | 389 | | 2 | 0050 | 805 |
| W: | ΑE, | AG, | AL, | AM, | ΑT, | ΑU, | ΑZ, | BA, | BB, | BG, | BR, | BW, | BY, | ΒZ, | CA, | CH, |
| | CN, | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EC, | EE, | EG, | ES, | FI, | GB, | GD, |
| | GE, | GH, | GM, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | KE, | KG, | KM, | ΚP, | KR, | ΚZ, |
| | LC, | LK, | LR, | LS, | LT, | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | MZ, | NA, |
| | NG, | NI, | NO, | ΝZ, | OM, | PG, | PH, | PL, | PT, | RO, | RU, | SC, | SD, | SE, | SG, | SK, |
| | SL, | SM, | SY, | ΤJ, | TM, | TN, | TR, | TT, | TZ, | UA, | UG, | US, | UΖ, | VC, | VN, | YU, |
| | ΖA, | ZM, | zw | | | | | | | | | | | | | |
| RW: | ΑT, | BE, | BG, | CH, | CY, | CZ, | DE, | DK, | EE, | ES, | FI, | FR, | GB, | GR, | HU, | ΙE, |
| | IS, | ΙT, | LT, | LU, | LV, | MC, | NL, | PL, | PT, | RO, | SE, | SI, | SK, | TR, | BF, | ВJ, |
| | CF, | CG, | CI, | CM, | GΑ, | GN, | GQ, | GW, | ML, | MR, | NE, | SN, | TD, | TG, | BW, | GH, |
| | GM, | KΕ, | LS, | MW, | MZ, | NA, | SD, | SL, | SZ, | TZ, | ŪĠ, | ZM, | ZW, | AM, | ΑZ, | BY, |
| | KG, | ΚZ, | MD, | RU, | ТJ, | TM | | | | | | | | | | |

PRIORITY APPLN. INFO.: JP 2004-233660 A 20040810 Disclosed is a powdery composition for transnasal administration which contains a nonpeptidic nonproteinaceous drug and crystalline cellulose masses having a specific mesh-size as a carrier therefor. This composition can exert an immediate action of the drug and a high absorbability. For example, morphine hydrochloride 65 mg and Avicel PH-F20 (crystalline cellulose) 135 mg were blended and nasally administered to monkeys for the determination of pharmacokinetic parameters of morphine.

53902-12-8, Tranilast IT

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (transnasal powder composition having immediate action and high absorbability)

53902-12-8 CAPLUS RN

Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) CN (CA INDEX NAME)

THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 14

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

CAPLUS COPYRIGHT 2007 ACS on STN L9 ANSWER 70 OF 98

ACCESSION NUMBER: 2006:148238 CAPLUS

DOCUMENT NUMBER: 144:239929

TITLE: Drug eluting stents made from crosslinked

biodegradable materials and drugs

INVENTOR(S): Sung, Hsing-Wen; Liang, Hsiang-Fa; Huang, Chin-Tsung;

Tu, Hosheng

PATENT ASSIGNEE(S): Taiwan

SOURCE: U.S. Pat. Appl. Publ., 46 pp., Cont.-in-part of U.S.

Ser. No. 916,170.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 12

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|------------------------|------|----------|-----------------|----|----------|
| | | | | - | |
| US 2006034885 | A1 | 20060216 | US 2004-929047 | | 20040827 |
| US 2005019404 | A1 | 20050127 | US 2004-916170 | | 20040811 |
| PRIORITY APPLN. INFO.: | * | | US 2004-916170 | A2 | 20040811 |
| | | | US 2003-610391 | A2 | 20030630 |
| | | | US 2003-518050P | P | 20031107 |
| | - | | US 2004-547935P | P | 20040226 |
| | • | | US 2004-565438P | Р | 20040426 |
| | | | US 2004-574501P | Р | 20040526 |
| | | | US 2004-585775P | Р | 20040706 |

OTHER SOURCE(S): MARPAT 144:239929

The present invention relates to a drug-loaded biodegradable stent and methods for treating vulnerable plaques of a patient comprising a plurality of layers or zones, each layer or zone comprising its own specific biodegrdn. rate and its specific drug loading characteristics. In one embodiment, the layers and zones are configured and arranged, in combination, radially, circumferentially and longitudinally. For example, a stent made from genipin-crosslinked chitosan was loaded with Taxol for controlled release of the antitumor agent.

IT 53902-12-8, Tranilast

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (drug eluting stents made from crosslinked biodegradable materials and drugs)

RN53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

L9 ANSWER 71 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:1262679 CAPLUS

DOCUMENT NUMBER: 143:472649

TITLE: Diarylalkanes as potent inhibitors of binuclear

enzymes

INVENTOR(S): Jia, Qi; Zhao, Ji-Fu

PATENT ASSIGNEE(S): Unigen Pharmaceuticals, Inc., USA

SOURCE: U.S. Pat. Appl. Publ., 39 pp.

CODEN: USXXCO

DOCUMENT TYPE:

LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| | PAT | CENT | NO. | | | KIN | D | DATE | | | APPL | ICAT | ION | NO. | | D. | ATE | |
|--------|------|-------|------|------|-----|-----------|-----|-------|----------|-----|------|-------|------|-----|-----|------|------|-----|
| | US | 2005 | 2670 | 47 | | A1 | - | 2005 | 1201 | | บร 2 | 005- | 1392 | 00 | | 2 | 0050 | 527 |
| | AU | 2005 | 2494 | 93 | | A1 | | 2005 | 1215 | | AU 2 | 005- | 2494 | 93 | | 2 | 0050 | 527 |
| | CA | 2567 | 801 | | | A1 | | 2005 | 1215 | | CA 2 | 005- | 2567 | 801 | | 2 | 0050 | 527 |
| | WO | 2005 | 1178 | 49 | | A1 | | 2005 | 1215 | | WO 2 | 005-1 | US18 | 884 | | 2 | 0050 | 527 |
| | | W: | ΑE, | AG, | ΑL, | AM, | ΑT, | AU, | ΑZ, | BA, | BB, | BG, | BR, | BW, | BY, | ΒZ, | CA, | CH, |
| | | | CN, | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EC, | EE, | EG, | ES, | FI, | GB, | GD, |
| | | | GE, | GH, | GM, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | KE, | KG, | KM, | ΚP, | KR, | ΚZ, |
| | • | | LC, | LK, | LR, | LS, | LT, | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | ΜZ, | NA, |
| | | | NG, | NI, | NO, | NZ, | OM, | PG, | PH, | PL, | PT, | RO, | RU, | SC, | SD, | SE, | SG, | SK, |
| | | | SL, | SM, | SY, | ТJ, | TM, | TN, | TR, | TT, | TZ, | UΑ, | UG, | US, | UΖ, | VC, | VN, | YU, |
| | | | ZA, | ZM, | ZW | | | | | | | | | | | | | |
| | | RW: | BW, | GH, | GM, | KΕ, | LS, | MW, | MZ, | NA, | SD, | SL, | SZ, | TZ, | UG, | ZM, | ZW, | AM, |
| | | | ΑZ, | BY, | KG, | ΚZ, | MD, | RU, | ΤJ, | TM, | ΑT, | ΒE, | BG, | CH, | CY, | CZ, | DE, | DK, |
| | | | EE, | ES, | FI, | FR, | GB, | GR, | ΗU, | ΙE, | IS, | IT, | LT, | LU, | MC, | NL, | PL, | PT, |
| | | | RO, | SE, | SI, | SK, | TR, | BF, | ВJ, | CF, | CG, | CI, | CM, | GA, | GN, | GQ, | GW, | ML, |
| | | | MR, | ΝĖ, | SN, | TD, | TG | | | | | | | | | | | |
| | ΕP | 1748 | 767 | | | A1 | | 2007 | 0207 | | EP 2 | 005- | 7621 | 86 | | 2 | 0050 | 527 |
| | | R: | ΑT, | BE, | BG, | CH, | CY, | CZ, | DE, | DK, | EE, | ES, | FI, | FR, | GB, | GR, | HU, | ΙE, |
| | | | IS, | ΙT, | LI, | LT, | LU, | MC, | NL, | ΡL, | PT, | RO, | SE, | SI, | SK, | TR | | |
| PRIOR | YTIS | APP: | LN. | INFO | . : | | | | | 1 | US 2 | 004- | 5755 | 99P | | P 2 | 0040 | 528 |
| | | | | | | | | | | 1 | WO 2 | 005-t | US18 | 884 | 1 | W 20 | 0050 | 527 |
| OMITHE | | TTDOT | (0) | | | 843 153 | - m | 7 4 2 | 4000 | | | | | | | | | |

OTHER SOURCE(S): MARPAT 143:472649

The present invention implements a strategy that combines an enzyme inhibition assay with a chemical dereplication process to identify active plant exts. and the particular compds.-diarylalkanes and/or diarylalkanels within those exts. that specifically inhibit binuclear enzyme function. Included in the present invention are compns. of matter comprised of one or more of diarylalkanes and/or diarylalkanels, which inhibit the activity of binuclear enzymes, particularly tyrosinase and which prevent melanin overprodn. The present invention also provides a method for inhibiting the activity of a binuclear enzyme, particularly tyrosinase and a method for preventing and treating diseases and conditions related to binuclear enzyme function. The present invention further includes a method for preventing and treating melanin overprodn. and diseases and conditions of the skin related thereto. The method for preventing and treating diseases and conditions related to binuclear enzyme function and melanin overprodn.

is comprised of administering to a host in need thereof an effective amount of a composition comprising one or more diarylalkanes and/or diarylalkanels synthesized and/or isolated from one or more plants together with a pharmaceutically acceptable carrier.

IT 1135-24-6, 3-Methoxy-4-hydroxycinnamic acid

RL: RCT (Reactant); RACT (Reactant or reagent)

(diarylalkanes as inhibitors of binuclear enzymes)

RN 1135-24-6 CAPLUS

CN 2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)- (9CI) (CA INDEX NAME)

L9 ANSWER 72 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2005:497471 CAPLUS

DOCUMENT NUMBER:

143:32422

TITLE:

Crosslinkable biological material and angiogenic agent

for promoting angiogenesis

INVENTOR(S):

Sung, Hsing-Wen; Liang, Huang-Chien; Tu, Hosheng

PATENT ASSIGNEE(S):

Taiwan

SOURCE:

U.S. Pat. Appl. Publ., 56 pp., Cont.-in-part of U.S.

Ser. No. 408,176.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 12

PATENT INFORMATION:

| PATENT 1 | | KIND | DATE | APPLICATION NO. | DATE |
|--------------|--------------------|----------|-----------|---------------------------------------|----------------------------|
| US 2005 | | | 20050609 | US 2004-827673 | 20040419 |
| WO 9819 | 718 | A1 | 19980514 | WO 1997-US20113 | 19971104 |
| RW: | | DE, DK | | FR, GB, GR, IE, IT, | |
| | 237 DE, FR, GB, | | 20021127 | EP 2002-19186 | 19971104 |
| US 6608 | 040 | B1 | 20030819 | US 2001-297808 | 20010927 |
| US 20020 | 091445 | A1 | 20020711 | US 2002-67130 | 20020204 |
| US 69984 | 118 | B2 B1 | 20030408 | US 2003-408176 | 20030407 |
| AU 20042 | 289270 | A1 | 20050526 | US 2003-408176 AU 2004-289270 | 20041105 |
| CA 25451 | 136 | A1 | 20050526 | CA 2004-2545136 | 20041105 |
| | | | | EP 2004-818654 GB, GR, IT, LI, LU, | |
| | IE, SI, FI, | RO, CY | , TR, BG, | CZ, EE, HU, PL, SK, | IS |
| PRIORITY APP | LN. INFO.: | | | US 1996-30701P | P 19961105 |
| | | | | WO 1997-US20113 | W 19971104 |
| | | | | US 2001-297808 | A2 20010927 |
| | | | • | US 2002-67130 US 2003-408176 | A2 20020204 A2 20030407 |
| | | | | IIS 2003-408176 | P 20030407 |
| | | | | US 2003-492874P US 2003-518050P | P 20031107 |
| | | | • | US 2003-526434P | P 20031202 |
| | | | | US 2004-547935P | P 20040226 |
| | | | | US 2004-552517P | P 20040312 |
| | | | | EP 1997-947356 | A3 19971104 |

Amethod for promoting angiogenesis in a patient comprising providing crosslinkable biol. solution to the target tissue, wherein the crosslinkable biol. solution is loaded with at least one angiogenic agent. In one embodiment, the at least one angiogenic agent is a non-protein factor selected from a group consisting of ginsenoside Rg1, ginsenoside Re, combination thereof and the like. In another embodiment, the crosslinkable biol. solution of the present invention is broadly defined in a form or phase of solution, paste, gel, suspension, colloid or plasma that may be solidifiable thereafter. For example, to increase pore sizes and porosities within test samples, the acellular pericardia were treated with acetic acid and collagenase. Subsequently, acellular tissues were fixed in a 0.05% genipin at 37° for 3 days. Genipin, as a crosslinking agent, was significantly less cytotoxic compared to glutaraldehyde used as a control.

IT 53902-12-8, Tranilast

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (biomaterial modified with composition containing angiogenic agent and crosslinker for promoting angiogenesis)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

L9 ANSWER 73 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2005:497233 CAPLUS

DOCUMENT NUMBER:

143:32417

TITLE:

Drug-eluting stent having collagen drug carrier

chemically treated with genipin

INVENTOR(S):

Sung, Hsing-Wen; Chen, Mei-Chin; Tu, Peter Y.; Tu,

Hosheng

PATENT ASSIGNEE(S):

Taiwan

SOURCE:

U.S. Pat. Appl. Publ., 35 pp., Cont.-in-part of U.S.

Ser. No. 717,162.

CODEN: USXXCO

DOCUMENT TYPE:

Patent English

LANGUAGE:

1: 12

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | KIND DATE | APPLICATION NO. | DATE |
|-----------------|-----------------|------------------------|------------------|
| | | | |
| US 2005123582 | A1 20050609 | US 2004-811413 | 20040326 |
| WO 9819718 | A1 19980514 | WO 1997-US20113 | 19971104 |
| W: CA, CN, JP, | US | | |
| RW: AT, BE, CH, | DE, DK, ES, FI, | FR, GB, GR, IE, IT, LU | , MC, NL, PT, SE |
| EP 1260237 | A1 20021127 | EP 2002-19186 | 19971104 |
| R: DE, FR, GB, | IT | | |
| US 6608040 | B1 20030819 | US 2001-297808 | 20010927 |
| US 6624138 | B1 20030923 | US 2002-211656 | 20020802 |

Ô

| US 20031 | .91071 | A1 | 2003 | 1009 | | | | | |
|---------------|-------------|-----|---------|------|-------|--------|---------|----|------------|
| US 20051 | .63818 | A1 | 2005 | 0728 | US | 2003- | 610391 | | 20030630 |
| AU 20042 | 89270 | A1 | 2005 | 0526 | AU | 2004- | 289270 | • | 20041105 |
| CA 25451 | .36 | A1 | 2005 | 0526 | CA | 2004- | 2545136 | | 20041105 |
| EP 16893 | 22 | A1 | 2006 | 0816 | EP | 2004- | 818654 | | 20041105 |
| R: | AT, BE, CH, | | | | | | | | E, MC, PT, |
| | IE, SI, FI, | RO, | CY, TR, | BG, | CZ, E | E, HU, | PL, SK, | IS | |
| PRIORITY APPI | N. INFO.: | | | | US | 1996- | 30701P | P | 19961105 |
| | | | | | WO | 1997- | US20113 | W | 19971104 |
| | | | | | | | | | 20010927 |
| | | | | | US | 2002- | 211656 | A2 | 20020802 |
| | | | | | | | 610391 | | 20030630 |
| | | | | | US | 2003- | 492874P | | |
| | | | | | US | 2003- | 518050P | P | 20031107 |
| | | | | | US | 2003- | 717162 | A2 | 20031119 |
| | | | | | US | 2004- | 547935P | _ | 20040226 |
| | | | | | US | 2004- | 552517P | P | 20040312 |
| | | | | | EP | 1997- | 947356 | A3 | 19971104 |
| | | | | | US | 2002- | 393565P | P | 20020702 |
| | | | | | US | 2004- | 565438P | P | 20040426 |
| | | | | | | | 574501P | P | 20040526 |
| | | | | | US | 2004- | 610391 | Α | 20040630 |
| | | | | | | | 585775P | P | 20040706 |
| | _, | | | | WO | 2004-1 | JS37217 | W | 20041105 |

OTHER SOURCE(S): MARPAT 143:32417

AB A method for treating vulnerable plaques of a patient, comprising: providing a biodegradable stent comprising a first supporting zone made of a first biodegradable material, wherein the supporting zone comprises at least a portion of continuous circumference of the stent; and a second therapeutic zone made of a second biodegradable material, wherein the therapeutic zone comprises at least one bioactive agent; delivering the biodegradable stent to the vulnerable plaques; orienting the therapeutic zone at about the luminal surface of the vulnerable plaque; and releasing the at least one bioactive agent for treating the vulnerable plaques.

IT 53902-12-8, Tranilast

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (drug-eluting stent having collagen drug carrier chemical treated with genipin)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

L9 ANSWER 74 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2005:140590 CAPLUS

DOCUMENT NUMBER:

142:225670

TITLE:

Composition for heart disease, its active ingredients,

method to prepare same and uses thereof

INVENTOR(S):

Yan, Xijun; Wu, Naifeng; Guo, Zhixin; Ye, Zhengliang;

Liu, Yan

PATENT ASSIGNEE(S):

Peop. Rep. China

SOURCE:

U.S. Pat. Appl. Publ., 16 pp.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

APPLICATION NO. DATE DATE PATENT NO. KIND ----------_____ ---------20050217 US 2004-903110 20040730 US 2005037094 Α1 US 2003-491466P PRIORITY APPLN. INFO.: P 20030731

This invention provides a composition for heart disease comprising exts. from raw herbs of 80.0-97.0% Radix salviae miltorrhizae, 1.0-19.0% Panax notoginseng and 0.1-1.0% borneol and its active ingredients. This invention also provides a method for preparing said composition and the active ingredients of the composition Finally, this invention provides various uses of said compns. and the active ingredients. Preparation of hydroalcoholic exts. of the above plants is described.

IT 158732-59-3, Salvianolic acid f

RL: NPO (Natural product occurrence); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); USES (Uses) (composition for heart disease, its active ingredients, method to prepare

same

and uses thereof)

RN 158732-59-3 CAPLUS

CN 2-Propenoic acid, 3-[2-[(1E)-2-(3,4-dihydroxyphenyl)ethenyl]-3,4-dihydroxyphenyl]-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L9 ANSWER 75 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:77978 CAPLUS

DOCUMENT NUMBER: 142:162660

TITLE: Biodegradable stent with crosslinked bioactive agent

for slow release

INVENTOR(S): Sung, Hsing-Wen; Chen, Mei-Chin; Tu, Peter Y.; Tu,

Hosheng

PATENT ASSIGNEE(S): Taiwan

SOURCE: U.S. Pat. Appl. Publ., 45 pp., Cont.-in-part of U.S.

Ser. No. 610,391.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 12

PATENT INFORMATION:

| PATENT NO. | KIND DATE | APPLICATION NO. | DATE |
|----------------|--------------------|-----------------------|-------------|
| | | | |
| US 2005019404 | A1 20050127 | US 2004-916170 | 20040811 |
| US 2005163818 | A1 20050728 | US 2003-610391 | 20030630 |
| US 2006034885 | A1 20060216 | US 2004-929047 | 20040827 |
| AU 2004289270 | A1 20050526 | AU 2004-289270 | 20041105 |
| CA 2545136 | A1 20050526 | CA 2004-2545136 | 20041105 |
| EP 1689322 | A1 20060816 | EP 2004-818654 | 20041105 |
| R: AT, BE, CH, | DE, DK, ES, FR, GB | , GR, IT, LI, LU, NL, | SE, MC, PT, |
| IE, SI, FI, | RO, CY, TR, BG, CZ | . EE. HU. PL. SK. IS | |

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US 2005163821
                                  20050728
                                               US 2005-906239
                                                                        20050210
                            A1
     WO 2006033686
                            A1
                                  20060330
                                               WO 2005-US19930
                                                                        20050608
             AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA,
             NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK,
              SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU,
              ZA, ZM, ZW
         RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
              IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF,
              CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM,
             KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG,
             KZ, MD, RU, TJ, TM
PRIORITY APPLN. INFO.:
                                               US 2003-610391
                                                                     A2 20030630
                                               US 2003-518050P
                                                                     P
                                                                        20031107
                                                                     Р
                                               US 2004-547935P
                                                                        20040226
                                               US 2004-565438P
                                                                     P
                                                                        20040426
                                                                     P
                                               US 2004-574501P
                                                                        20040526
                                                                     P
                                               US 2004-585775P
                                                                        20040706
                                                                     Р
                                               US 1996-30701P
                                                                        19961105
                                                                     W
                                                                        19971104
                                               WO 1997-US20113
                                                                     A2 20010927
                                               US 2001-297808
                                               US 2002-211656
                                                                     A2 20020802
                                               US 2004-610391
                                                                     Α
                                                                        20040630
                                               US 2004-916170
                                                                     A2 20040811
                                               WO 2004-US37217
                                                                     W
                                                                        20041105
                                               US 2004-24101
                                                                     A2 20041228
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OTHER SOURCE(S): MARPAT 142:162660

AB The present invention relates to a drug-loaded biodegradable stent or implant for drug slow release and methods for treating vulnerable plaques of a patient comprising a plurality of layers or zones, each layer or zone comprising its own specific biodegrdn. rate and its specific drug loading characteristics. Specifically, the layers and zones are configured and arranged, in combination, radially, circumferentially and longitudinally. The crosslinked biodegradable stent or implant comprises at least one layer or zone of biol. material, said biol. material comprising at least one bioactive agent and being crosslinked with a means for crosslinking said biol. material.

IT 53902-12-8, Tranilast

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (biodegradable stent with crosslinked bioactive agent for slow release) 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

L9 ANSWER 76 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2004:995641 CAPLUS

DOCUMENT NUMBER:

141:416007

TITLE:

RN

Pharmaceutical compositions containing drugs entrapped

in crosslinked ionic core micelles

INVENTOR(S):
PATENT ASSIGNEE(S):

Bronich, Tatiana K.; Kabanov, Alexander V. University of Nebraska Board of Regents, USA

SOURCE: U.S. Pat. Appl. Publ., 17 pp.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|---------------------|---------|--------------|-------------------------|---------------|
| | | | | | |
| | US 2004228823 | A1 | 20041118 | US 2003-440221 | 20030516 |
| | RITY APPLN. INFO.: | | | | 20030516 |
| AB | | | | micelles with cross-li | |
| | cores as delivery v | ehicles | for therape | utics, diagnostics, nuc | leic acids, |
| | proteins, small mol | s. and | the like. T | he present invention pr | ovides addnl. |
| | methods of synthesi | s and u | ses for such | micelles. For example | , cisplatin |
| | was entrapped into | the mic | elles of eth | ylene oxide-sodium meth | acrylate |
| | block copolymer com | plexed | with calcium | ion and crosslinked by | 1,2-ethylene |
| | diamine. | _ | | <u>-</u> | - |

1135-24-6, 4-Hydroxy-3-methoxy cinnamic acid ΙT

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (polyanionic segment; pharmaceutical micelles containing drugs entrapped in crosslinked ionic core micelles)

RN 1135-24-6 CAPLUS

2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)- (9CI) (CA INDEX NAME) CN

L9 ANSWER 77 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2001:676586 CAPLUS

DOCUMENT NUMBER:

135:216027

TITLE:

Stretchable patches comprising drugs in tacky layers

INVENTOR(S):

Hidaka, Osafumi; Ohata, Akiko

PATENT ASSIGNEE(S):

Teijin Limited, Japan; Teysan Pharmaceuticals Co.,

Ltd.

SOURCE:

PCT Int. Appl., 24 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PA | PATENT NO. | | | | KIND DATE | | | | APPLICATION NO. | | | | | DATE | | | |
|-----|------------|----------|--------|-----|------------|-------------|------|------|-----------------|----------------|------|------|-----|------|----------|-------|-----|
| WO. | 2001 | 0660 | 95 | | Δ1 | 71 20010012 | | | 1 | WO 2001-JP1691 | | | | | 20010305 | | |
| | | | | | | | | | | | | | | | | | |
| | w: | | | | | | | | | | | | | | | CH, | |
| | | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DΖ, | EE, | ES, | FΙ, | GB, | GD, | GE, | GH, | GM, |
| | | HR, | HU, | ID, | IL, | IN, | IS, | JP, | ΚE, | KG, | KΡ, | KR, | ΚZ, | LC, | LK, | LR, | LS, |
| | | LT, | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | MZ, | NO, | NZ, | PL, | PT, | RO, |
| | | RU, | SD, | SE, | SG, | SI, | SK, | SL, | ТJ, | TM, | TR, | TT, | TZ, | UΑ, | ŪĠ, | US, | UΖ, |
| | | VN, | YU, | ZA, | ZW, | AM, | ΑZ, | BY, | KG, | ΚZ, | MD, | RU, | ТJ, | TM | | | |
| | RW: | GH, | GM, | KE, | LS, | MW, | ΜZ, | SD, | SL, | SZ, | TZ, | ŬĠ, | ZW, | AT, | BE, | CH, | CY, |
| | | DE, | DK, | ES, | FI, | FR, | GB, | GR, | ΙE, | IT, | LU, | MC, | NL, | PT, | SE, | TR, | BF, |
| | | ВJ, | CF, | CG, | CI, | CM, | GΑ, | GN, | GW, | ML, | MR, | ΝE, | SN, | TD, | TG | | |
| CA | 2373 | 171 | | | A 1 | : | 2001 | 0913 | (| CA 2 | 001- | 2373 | 171 | | 2 | 00103 | 305 |
| ΑU | 2001 | 0360 | 92 | | A 5 | : | 2001 | 0917 | 1 | AU 2 | 001- | 3609 | 2 | | 2 | 0010 | 305 |
| ΑU | 7808 | 81 | | | B2 | | 2005 | 0421 | | | | | | | | | |

EP 1177786 Α1 20020206 EP 2001-908314 20010305 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,

IE, SI, LT, LV, FI, RO

US 2002146445 US 2001-959420 A1 20021010 20011026 PRIORITY APPLN. INFO.: JP 2000-61676 20000307 Α WO 2001-JP1691 W 20010305

This invention relates to a stretchable patch which comprises a support AB film having a thickness of 1 to 50 µm and a drug-containing tacky layer having a thickness of 3 to 400 µm, wherein the support film satisfies the following requirements (1) to (4): (1) the support film comprises a copolymer obtained by copolymg. 0-90 % vinyl acetate, 10-97 % alkyl (meth) acrylate in which the alkyl has 3 to 14 carbon atoms on the average, and 0-15 % (meth)acrylic acid; (2) the copolymer has a degree of crosslinking of 20 or higher when crosslinked with a polyvalent metal, and when the copolymer is crosslinked with a polyfunctional chain compound, the content of units derived from the compound in the copolymer is 1 to 10; (3) the support film has a strength of self-adhesion of 150 g or lower; and (4) the support film has an elastic recovery at 10 elongation of 70 or higher.

53902-12-8, Tranilast IT

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (stretchable patches containing drugs in tacky layers)

53902-12-8 CAPLUS RN

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

REFERENCE COUNT: THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS 1 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

T.9 ANSWER 78 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

1998:365197 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 129:81875

TITLE: Certain Norditerpenoid Alkaloids and Their

Cardiovascular Action

AUTHOR (S): Desai, Haridutt K.; Hart, Bradley P.; Caldwell, R.

William; Huang, Jianzhong; Pelletier, S. William

CORPORATE SOURCE: Institute for Natural Products Research and Department

of Chemistry, University of Georgia, Athens, GA,

30602-2556, USA

SOURCE: Journal of Natural Products (1998), 61(6), 743-748

CODEN: JNPRDF; ISSN: 0163-3864

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal LANGUAGE: English

Thirteen new derivs. of norditerpenoid alkaloids, namely, 8-deacetyl-8-p-aminobenzoyldelphinine, 8-deacetyl-8-anthranoyldelphinine,

8-deacetyl-8-(4-hydroxy-3-methoxycinnamoyl)delphinine,

16-demethoxy-15,16-didehydro-8-p-anisoyl-14-benzoyldelphonine,

6-acetylheteratisine N-oxide, 3,8-diacetylfalconerine,

8-stearoylfalconerine, 8-linolenylfalconerine, 13-acetylpyrodelphinine,

13-acetyldelphinine N-oxide, N-deacetyl-8,9-diacetyllappaconitine,

8,9-(methylenedioxy)lappaconine, and 16-epipyroaconitine N-oxide, were prepared, and their structures were established by anal. of spectroscopic data (1D and 2D NMR, HRFABMS). The preliminary in vivo cardiovascular action (hypotensive, bradycardic, and ventricular arrhythmias) of these

new compds. was tested in male Sprague-Dawley rats. The results are reported herein.

IT 537-98-4, trans-4-Hydroxy-3-methoxycinnamic acid

RL: RCT (Reactant); RACT (Reactant or reagent)
(preparation of norditerpenoid alkaloids as cardiovascular agents)

RN 537-98-4 CAPLUS

CN 2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)-, (2E)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

REFERENCE COUNT: 12 THERE ARE 12 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 79 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1996:209937 CAPLUS

DOCUMENT NUMBER: 124:242363

TITLE: Stable pharmaceutical lipid emulsions containing oils

and emulsifiers and lecithins

INVENTOR(S): Suzuki, Hidekazu; Yamazaki, Satoshi; Naito, Yoshikazu;

Endo, Kenji; Oguma, Touru; Maeda, Makoto

APPLICATION NO.

DATE

PATENT ASSIGNEE(S): Wakamoto Pharmaceutical Co., Ltd., Japan

' KIND

DATE

SOURCE: Can. Pat. Appl., 77 pp.

CODEN: CPXXEB

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.

| CA 2153553 | | 19960114 | | |
|--------------------|---------------|-------------|-----------------------|------------------------|
| US 5693337 | | 19971202 | | |
| EP 700678 | A1 | | EP 1995-110923 | |
| | R, GB, IT | 233,00023 | 21 1333 110323 | 1,750,12 |
| JP 08081360 | A A | 19960326 | JP 1995-197896 | 19950712 |
| PRIORITY APPLN. IN | = = | 13300320 | JP 1994-183045 | |
| | - • | prices (A) | an oil component, (| |
| | | | | |
| | | | | and (C) water, wherein |
| | | | citric acid or a pha | |
| | | | one member selected | |
| | | | ine, serine, histidi: | |
| | | | ereof, provided that | |
| simultaneousl | y contain met | hionine and | d phenylalanine. The | e emulsion does |
| not change of | color and fo | rmation of | oil drops associated | d with the |
| conventional : | natural lecit | hin-contair | ning lipid emulsions | due to the |
| | | | additives. The drug | |
| | | | stability and thus | |
| | | | gs such as injections | |
| | | | halants and drugs for | |
| | | | numectants. A soluti | |
| | • | | vas added to a solut: | |
| | | | | |
| | | | g of yolk lecithin | |
| | | | collowed by evaporat: | |
| obtain a lipid | d film. To t | he lipid fi | ilm was added 5.4 g o | of soybean oil and |

94 mL of 2% glycerin aqueous solution followed by vigorous stirring through shaking to carry out preliminary emulsification. The preliminarily emulsified liquid was passed through microfluidizer 10 times under a pressure of 750 kg/cm2 to emulsify the liquid, the pH value of the emulsified liquid was adjusted to 6.5-7.5 to give a milk white stock lipid emulsion.

IT 53902-12-8, Tranilast

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(stable pharmaceutical lipid emulsions containing oils and emulsifiers and lecithins)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

L9 ANSWER 80 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1986:116113 CAPLUS

DOCUMENT NUMBER: 104:116113

TITLE: Lipid nanopellet oral drug formulation

INVENTOR(S): Speiser, Peter

PATENT ASSIGNEE(S): Rentschler, Dr., Arzneimittel G.m.b.H. und Co., Fed.

Rep. Ger.

SOURCE: Ger. Offen., 35 pp.

CODEN: GWXXBX

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--------|--------------|-----------------|-------------|
| | | | | |
| DE 3421468 | A1 | 19851219 | DE 1984-3421468 | 19840608 |
| EP 167825 | A2 | 19860115 | EP 1985-106926 | 19850604 |
| EP 167825 | A3 | 19870121 | | |
| EP 167825 | B1 | 19900808 | | |
| R: AT, BE, CH, | DE, FR | , GB, IT, LI | , LU, NL, SE | |
| AT 55243 | T | 19900815 | AT 1985-106926 | 19850604 |
| JP 61056122 | Α | 19860320 | JP 1985-120726 | 19850605 |
| US 4880634 | Α | 19891114 | US 1987-66459 | 19870626 |
| PRIORITY APPLN. INFO.: | | | DE 1984-3421468 | A 19840608 |
| | • | | EP 1985-106926 | A 19850604 |
| | | | US 1985-740771 | A1 19850630 |
| | | | | |

AB Lipid nanopellets (80-800 nm), as aqueous colloidal suspensions, are carrier systems for oral drugs. The lipids are saturated fatty acids, their esters with glycerol and with other polyalcs., and fatty alcs. The system contains natural or artificial surfactants. Thus, a mixture of 2 g tristearin and 0.6 g testosterone undecanoate was melted at 85° and 0.4 g phospholipon 100-H in 4 mL CHCl3 was added. The CHCl3 was evaporated and 0.04 Na cholate in 200 mL water was added, followed by stirring and ultrasonication, to give the nanopellet suspension.

IT 53902-12-8

RN

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (lipid nanopellets, for oral administration as aqueous colloidal emulsion) 53902-12-8 CAPLUS

CAPLUS COPYRIGHT 2007 ACS on STN ANSWER 81 OF 98

1986:28398 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 104:28398

Derivatives of isoquinoline. XXIV. Synthesis and TITLE:

> biological properties of 1- and 2-arylalkenyl-6,7dimethoxy-4,4-diethyl-substituted hydrogenated derivatives of isoquinoline and their noncyclic

analogs

AUTHOR (S): Airapetyan, G. K.; Avetisyan, A. S.; Markaryan, E. A.;

Pogosyan, A. V.

CORPORATE SOURCE: USSR

SOURCE: Deposited Doc. (1984), VINITI 3943-84, 10 pp. Avail.:

VINITI

DOCUMENT TYPE: Report

LANGUAGE: Russian

GI

AB Twenty title compds. were prepared (1) by condensation of the chloranhydrides of α,β -unsatd. acids with 6,7-dimethoxy-4,4diethyl-1,2,3,4-tetrahydroisoquinoline [72193-99-8] to produce cyclic 1-arylalkenyls [I; X = O or H2; R = H, Me, or Ph; R1 = Ph or (MeO) 2C6H3], (2) by condensation with 2-(3,4-dimethoxyphenyl)-2-ethylbutylamine [99611-90-2] to produce acyclic analogs [II; R = H, Me, or Ph; R1 = Ph or (MeO)2C6H3], or (3) by cyclization of II to produce 2-arylalkenyls [III; R = H or Ph; R1 = Ph or (MeO)2C6H3]. The hydrogenated derivs. of I, II, and III were prepared by reduction with AlH3 or LiAlH4. Hydrochlorides of some of the compds. were also prepared Studies in isolated rat vas deferens prepns. showed that several hydrochlorides of hydrogenates I and III possessed weak sympatholytic activity. At dosages of 0.1-3.0 mg/kg in unspecified laboratory animals, none of the compds. affected systemic arterial pressure, with the exception of a hydrogenated methylphenyl derivative of III [99611-97-9], which exhibited marked hypotensive activity.

IT 99612-04-1P

> RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses)

(preparation and pharmacol. of)

RN99612-04-1 CAPLUS

2-Propenamide, 3-(3,4-dimethoxyphenyl)-N-[1-(3,4-dimethoxyphenyl)-1-CNethylpropyl] - (9CI) (CA INDEX NAME)

CAPLUS COPYRIGHT 2007 ACS on STN L9 ANSWER 82 OF 98

ACCESSION NUMBER: 1976:577752 CAPLUS

DOCUMENT NUMBER: 85:177752

3-Isorescinnamine derivatives TITLE:

INVENTOR(S): Tanaka, Yoshihiro; Fujimoto, Yasuo Nippon Chemiphar Co., Ltd., Japan PATENT ASSIGNEE(S):

SOURCE: Jpn. Kokai Tokkyo Koho, 4 pp.

CODEN: JKXXAF

DOCUMENT TYPE:

Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|------------------------|------|----------|-----------------|---|----------|
| | | | | | |
| JP 51019798 | Α | 19760217 | JP 1974-89782 | | 19740807 |
| PRIORITY APPLN. INFO.: | | | JP 1974-89782 | A | 19740807 |

$$R^{20}$$
 R^{40}
 R^{30}
 R^{10}
 R^{40}
 R^{4

AB 3-Isorescinnamine derivs. I (R1 to R5 = lower alkyl) were prepared by reaction of alkyl 3-isoreserpate or its derivs. (II) with cinnamic acids III or their functional derivs. I had hypotensive action (no data). Thus, 5.0 g II (R1 = R2 = R3 = Me) in DMF-C5H5N was treated with 10.5 g

III (R4 = Me, R5 = Et) at room temperature to give, after treatment with HNO3, 6.4 g I.HNO3 (R1 = R2 = R3 = R4 = Me, R5 = Et) (IV). Free IV was also

prepared 42381-67-9

TΤ

RL: RCT (Reactant); RACT (Reactant or reagent)

(acylation of isoreserpate by)

RN 42381-67-9 CAPLUS

CN 2-Propenoic acid, 3-[4-[(ethoxycarbonyl)oxy]-3-methoxyphenyl]- (9CI) (CA INDEX NAME)

L9 ANSWER 83 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1976:180461 CAPLUS

DOCUMENT NUMBER:

84:180461 Rescinnamines

TITLE: INVENTOR(S):

Kametani, Tetsuji

PATENT ASSIGNEE(S):

Japan Chemipha Co., Ltd., Japan

SOURCE:

Jpn. Kokai Tokkyo Koho, 3 pp. Division of Japan. Kokai

73 40,800.

CODEN: JKXXAF

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|------------------------|------|----------|-----------------|---|----------|
| | | | | - | |
| JP 50123699 | A | 19750929 | JP 1974-64929 | | 19740610 |
| PRIORITY APPLN. INFO.: | | | JP 1974-64929 | Α | 19740610 |
| GI | | | | | |

$$R^{10}$$
 H
 H
 $R^{2}O_{2}C$
 O_{R}^{3}
 R^{2}
 R^{2}
 R^{2}
 R^{2}
 R^{2}
 R^{2}
 R^{2}
 R^{2}
 R^{2}

AB Rescinnamines I [R1-3 = alkyl; R4 = alkoxy, R5 = (alkoxycarbonyl)oxy, R6 = R7 = H; or R4 = R5 = R6 = alkoxy, R7 = NO2] were prepared by acylating alkyl reserpates with the appropriate cinnamic acids or their reactive derivs. I have hypotensive effect (no data). Thus, 3-[(ethoxycarbonyl)oxy]-4-methoxycinnamic acid was heated with SOCl2, and then treated with Me reserpate in C5H5N to give 75% I (R1 = R2 = R3 = Me, R4 = 4-MeO, R5 = 3-EtO2CO, R6 = R7 = H). Also prepared were I (R1 = R2 = R3 = Me) (R4, R5, R6, R7 given): 3-MeO, 4-EtO2CO, H, H; 3,4,5-(MeO)3, NO2.

IT 59189-24-1

RL: RCT (Reactant); RACT (Reactant or reagent) (acyl chlorination of)

RN59189-24-1 CAPLUS

2-Propenoic acid, 3-[3-[(ethoxycarbonyl)oxy]-4-methoxyphenyl]- (9CI) (CA CN INDEX NAME)

ANSWER 84 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1974:403945 CAPLUS

DOCUMENT NUMBER: 81:3945

N1-2-Imidazolinyl carbohydrazides TITLE:

PATENT ASSIGNEE(S): Ferlux-Chimie S. A. SOURCE: Fr. Demande, 20 pp.

CODEN: FRXXBL

DOCUMENT TYPE: Patent LANGUAGE: French

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--------|--------------|-----------------|----------|
| | | | | |
| FR 2186238 | A1 | 19740111 | FR 1972-16056 | 19720505 |
| FR 2186238 | B1 | 19750620 | | |
| PRIORITY APPLN. INFO.: | | • | FR 1972-16056 A | 19720505 |
| GI For diagram(s), see | printe | ed CA Issue. | | |

AB Hydrazides I (R = aryloxyalkl, arylvinyl, alkyl, aryl, 4-pyridyl) (41 compds.) were prepared, e.g. by acylating 2-hydrazino-2-imidazoline. I [R = 2,4-MeO(CH2:CHCH2)C6H3OCH2] at 80 mg/kg orally in mice gave 50% protection in the writhing syndrome test, and at 20 mg/kg i.v. in rats showed a strong, persistent hypotensive effect.

52377-39-6P IT

L9

RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of)

52377-39-6 CAPLUS RN

2-Propenoic acid, 3-(3,4,5-trimethoxyphenyl)-, 2-(4,5-dihydro-1H-imidazol-CN2-yl)hydrazide, monohydrochloride (9CI) (CA INDEX NAME)

HCl

ACCESSION NUMBER: 2006:34388 CAPLUS

DOCUMENT NUMBER: 144:114303

TITLE: Extraction of sweet potato

INVENTOR(S): Takagaki, Kinya

PATENT ASSIGNEE(S): Toyo Shinyaku Co., Ltd., Japan SOURCE: Jpn. Kokai Tokkyo Koho, 17 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|----------|------------|---------------------|------------------|
| | | | | |
| JP 2006008665 | A | 20060112 | JP 2005-147373 | 20050519 |
| PRIORITY APPLN. INFO.: | | | JP 2004-152599 | A 20040521 |
| AB This invention pro- | vides an | extraction | method of sweet pot | ato to obtain an |
| extract | | | | |

with higher content. of polyphenols. Sweet potato stems and leaves are extracted using water or water-containing organic solvents, preferably an aqueous

ethanolic solution The exts. comprise tricaffeoylquinic acid and/or dicaffeoylquinic acid. The exts. may be used as antidiabetics, antihypertensives, etc. (no data given).

IT 71275-40-6P

RL: NPO (Natural product occurrence); PUR (Purification or recovery); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation) (extraction of sweet potato)

RN 71275-40-6 CAPLUS

CN Cyclohexanecarboxylic acid, bis[[3-(3,4-dihydroxyphenyl)-1-oxo-2-propenyl]oxy]dihydroxy-, $(1\alpha,3\alpha,4\alpha,5\beta)$ - (9CI) (CA INDEX NAME)

CM 1

CRN 36413-60-2 CMF C7 H12 O6

Relative stereochemistry.

CM 2

CRN 331-39-5 CMF C9 H8 O4

HO OH CH
$$=$$
 CH $-$ CO₂H

ANSWER 86 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN L9

2005:64670 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 142:254230

Tranilast attenuates cardiac matrix deposition in TITLE:

experimental diabetes: role of transforming growth

factor-β

Martin, Jennifer; Kelly, Darren J.; Mifsud, Sally A.; AUTHOR (S):

Zhang, Yuan; Cox, Alison J.; See, Fiona; Krum, Henry;

Wilkinson-Berka, Jennifer; Gilbert, Richard E.

University of Melbourne Department of Medicine, St. CORPORATE SOURCE:

Vincent's Hospital, Australia

Cardiovascular Research (2005), 65(3), 694-701 SOURCE:

CODEN: CVREAU; ISSN: 0008-6363

Elsevier B.V. PUBLISHER:

DOCUMENT TYPE: Journal English LANGUAGE:

The pathol. accumulation of extracellular matrix is a characteristic feature of diabetic cardiomyopathy that is directly related to a loss of function. Tranilast [n-(3,4-anthranilic acid)], used for the treatment of fibrotic skin diseases, has also been shown to inhibit transforming growth factor- β (TGF- β)-induced matrix production in kidney epithelial To investigate the effects of tranilast in the diabetic heart, we examined its effects in cultured cardiac fibroblasts and then assessed its effects in (mRen-2)27 diabetic rats with established disease (8 wk after streptozotocin). In vitro studies demonstrated a 58% reduction in TGF-β1-induced 3[H]-hydroxyproline incorporation with tranilast 30 μΜ (p<0.01). At 16 wk, diabetes in the Ren-2 rat was associated with increased cardiac fibrosis and evidence of TGF-β1 activation, as measured by the abundance of phosphorylated Smad2. Despite persistent hyperglycemia and hypertension, tranilast attenuated cardiac fibrosis by 37% (p<0.05) in association with reduction in phospho-Smad2

(p<0.01).

These findings indicate that tranilast has antifibrotic actions in the Ren-2 model of exptl. diabetic cardiac disease by mechanisms that might attributable to reduced TGF-β activity.

53902-12-8, Tranilast IT

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(tranilast attenuates cardiac matrix deposition in exptl. diabetes and role of TGF-β)

53902-12-8 CAPLUS RN

Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) CN (CA INDEX NAME)

THERE ARE 38 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 38 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 87 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2004:777463 CAPLUS

DOCUMENT NUMBER: 142:127070

TITLE: Tranilast Attenuates Structural and Functional Aspects

of Renal Injury in the Remnant Kidney Model

AUTHOR(S): Kelly, Darren J.; Zhang, Yuan; Gow, Renae; Gilbert, Richard E.

CORPORATE SOURCE: Departments of Medicine, St. Vincent's Hospital,

University of Melbourne, Australia

Journal of the American Society of Nephrology (2004), SOURCE:

15(10), 2619-2629

CODEN: JASNEU; ISSN: 1046-6673 Lippincott Williams & Wilkins

PUBLISHER: DOCUMENT TYPE: Journal English LANGUAGE:

Pathol. fibrosis is a key feature of progressive renal disease that AΒ correlates closely with kidney dysfunction and in which the prosclerotic growth factor TGF-eta has been consistently implicated. Tranilast (n-[3,4-dimethoxycinnamoyl] anthranilic acid), an antifibrotic agent that is used to treat hypertrophic scars and scleroderma, has also been shown to inhibit TGF-eta-induced extracellular matrix synthesis in a range of cell types, including those of renal origin. Therefore, the effects of tranilast on kidney fibrosis and dysfunction were examined in the subtotal nephrectomy model of progressive renal injury. Subtotal nephrectomy led to proteinuria and renal dysfunction in association with glomerulosclerosis, tubulointerstitial fibrosis, and macrophage accumulation. Despite persistent hypertension, treatment with tranilast led to a reduction in albuminuria (61.7 +/ \div 1.2 vs. 20.5 +/ \div 1.3 mg/d; P < 0.01) and plasma creatinine (0.16 vs. 0.08 mmol/L; P < 0.01) in subtotally nephrectomized rats. In addition, features suggestive of $TGF-\beta$ activation, including glomerulosclerosis, tubulointerstitial fibrosis, tubular atrophy, and macrophage accumulation, all were significantly attenuated by tranilast in association with evidence of reduced TGF- β signaling in vivo. In the context of a recent pilot study in humans, the findings of the present report suggest that tranilast may provide a novel strategy for the treatment of progressive kidney disease characterized by fibrotic scarring.

IT 53902-12-8, Tranilast

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(tranilast attenuated structural, functional aspects of renal disease with reduced glomerulosclerosis, tubulointerstitial fibrosis, tubular atrophy, macrophage accumulation, proteinuria, creatinine clearance in subtotally nephrectomized rat)

₽N 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

THERE ARE 50 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 50 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 CAPLUS COPYRIGHT 2007 ACS on STN ANSWER 88 OF 98

ACCESSION NUMBER: 1996:38949 CAPLUS

DOCUMENT NUMBER: 124:76537

TITLE: Cardiac hypertrophy inhibitors containing tranilast INVENTOR (S): Nakajima, Mitsuyoshi; Umemura, Kazuo; Kikuchi, Shinji

PATENT ASSIGNEE(S): Kissei Pharmaceutical, Japan SOURCE: Jpn. Kokai Tokkyo Koho, 4 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

APPLICATION NO. PATENT NO. KIND DATE DATE ______ ______ _ _ _ _ **---**---_____ JP 1994-105882 19951024 JP 07277966 Α 19940408 JP 1994-105882 PRIORITY APPLN. INFO.:

B Prophylactic and therapeutic agents for cardiac hypertrophy contain 2-(3,4-dimethoxycinnamoyl)aminobenzoic acid (I) or its pharmacol. acceptable salts as an active ingredient. Administration of I at 300 mg/kg to spontaneously hypertensive rats once a day for 4 wk suppressed hypertrophy of left ventricle.

IT 53902-12-8, Tranilast

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(cardiac hypertrophy inhibitors containing tranilast)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

L9 ANSWER 89 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1996:30299 CAPLUS

DOCUMENT NUMBER:

124:135194

TITLE:

Tranilast suppresses intimal hyperplasia after

photochemically induced endothelial injury in the rat

AUTHOR(S):

Kikuchi, Shinji; Umemura, Kazuo; Kondo, Kazunao;

Nakashima, Mitsuyoshi

CORPORATE SOURCE:

Hamamatsu, 431-31, Japan

SOURCE:

European Journal of Pharmacology (1996), 295(2/3),

221-7

CODEN: EJPHAZ; ISSN: 0014-2999

PUBLISHER: Elsevier DOCUMENT TYPE: Journal LANGUAGE: English

AB Intimal thickening in the femoral artery of spontaneously hypertensive rats (SHR) was initiated by endothelial damage induced by the photochem. reaction between green light and systemic rose bengal. This model represents a non-mech. method of producing vessel wall denudation. Neointima formation was assessed by calculating the cross-sectional area of intima, media and lumen, using computer anal. Tranilast (30, 100 and 300 mg/kg, p.o.), administered 2 days prior to endothelial injury, reduced intimal area by 29, 62 and 87%, resp., compared to that of vehicle-treated controls. In cultured SHR-derived vascular smooth muscle cells, tranilast produced concentration-dependent inhibition of mitogenesis, whether stimulated by platelet-derived growth factor, basic fibroblast growth factor, insulin-like growth factor or fetal bovine serum. These results suggest that tranilast may be effective in preventing coronary restenosis.

IT 53902-12-8, Tranilast.

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES

(Uses)

(tranilast suppresses intimal hyperplasia after photochem. induced endothelial injury in rat)

RN 53902-12-8 CAPLUS

Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) CN (CA INDEX NAME)

ANSWER 90 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1995:883416 CAPLUS

DOCUMENT NUMBER: 124:80349

Kukoamine A and other hydrophobic acylpolyamines: TITLE:

potent and selective inhibitors of Crithidia

fasciculata trypanothione reductase

AUTHOR(S): Ponasik, James A.; Strickland, Corey; Faerman, Carlos;

Savvides, Savvas; Karplus, P. Andrew; Ganem, Bruce

CORPORATE SOURCE: Dep. Chem., Cornell Univ., Ithaca, NY, 14853-1301, USA SOURCE: Biochemical Journal (1995), 311(2), 371-5

CODEN: BIJOAK; ISSN: 0264-6021

PUBLISHER: Portland Press

DOCUMENT TYPE: Journal LANGUAGE: English

The enzyme trypanothione reductase (TR), together with its substrate, the glutathione-spermidine conjugate trypanothione, plays an essential role in protecting parasitic trypanosomatids against oxidative stress and is a target for drug design. Here the authors show that a naturally occurring spermine derivative, the antihypertensive agent kukoamine A [N1N12-bis(dihydrocaffeoyl)-spermine] inhibits TR as a mixed inhibitor (K1 = 1.8 μM , Kii = 13 μM). Kukoamine shows no significant inhibition of human glutathione reductase (Ki > 10 mM) and thus provides a novel selective drug lead. The corresponding N1N8-bis(dihydrocaffeoyl)spermidin e derivative was synthesized and acted as a purely competitive inhibitor with Ki = 7.5 μM. A series of mono- and di-acylated spermines and spermidines were synthesized to gain an insight into the effect of polyamine chain length, the nature and position of the acyl substituent and the importance of conformational mobility. These compds. inhibited TR with Ki values ranging from 11 to 607 μM.

IT 171294-45-4

CN

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)

(kukoamine A and other hydrophobic acylpolyamines as potent and selective inhibitors of Crithidia fasciculata trypanothione reductase)

ŔŊ 171294-45-4 CAPLUS

> 2-Propenamide, N,N'-[1,4-butanediylbis(imino-3,1-propanediyl)]bis[3-(3,4dihydroxyphenyl) - (9CI) (CA INDEX NAME)

> > PAGE 1-A

L9 ANSWER 91 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1964:34188 CAPLUS

DOCUMENT NUMBER: 60:34188

ORIGINAL REFERENCE NO.: 60:6114e-h,6115a

TITLE: Urinary excretion of phenolic and indolic acids after

ingestion of α -methyldopa

AUTHOR(S): Ruge, W.; Hartmann, F.

CORPORATE SOURCE: Univ. Marburg a.d. Lahn, Germany

SOURCE: Klinische Wochenschrift (1963), 41(17), 856-61

CODEN: KLWOAZ; ISSN: 0023-2173

DOCUMENT TYPE: Journal LANGUAGE: Unavailable

AB cf. CA 58, 1801h. By means of 2-dimensional paper chromatog. the urinary excretion pattern of phenolic and indolic acids was studied in 10 female albino rats (Siv 50 strain), and in a man and a woman who showed essential hypertension, before and after ingestion of α-methyldopa, Aldomet, (L-α-methyl-3,4-dihydroxy-1-phenylalanine) (I). Each rat

received 100 mg. I/day in drinking H2O. Urine samples for chromatog. were prepared by acidification and solvent extraction (EtOAc, 95% EtOH). The spot

put

on the paper represented urine containing 2.5 mg. creatinine. 4-Hydroxy-3-methoxymandelic acid (II) separated better if the solvent for the 1st dimension was iso-PrOH-NH3-H2O (80:2:18), and for the 2nd, PhH-HOAc-H2O (125:72:3). The phenolic acid spots were then developed with diazotized nitroaniline. For the indolic acids, the solvent for the 1st dimension was iso-PrOH-NH3-H2O (200:10:20), and for the 2nd, BuOH-HOAc-H2O (150:30:50) with Ehrlich's reagent as developer. During the tests on the 2 patients they were allowed neither coffee nor bananas, and were given daily 0.75-2.0 g. I according to tolerance. Blood pressures changed, resp., from 220/115 to 160/95 and from 180/110 to 200/120 during the 3 wk of tests wherein 1 control and 6 treated 24-h. urines were collected from each patient. On these there were quant. determined both 5-hydroxyindoleacetic acid (III) and II (Sandler and Ruthwen, CA 54, 16521f; 55, 23644h). Thus, in normal rat urine 13 different known phenolic acids were proved present, and after I was given, o-hydroxyhippuric and o-hydroxyphenylpyruvic (IV) acids also appeared; ferulic, vanillic (V), and p-hydroxybenzoic (VI) acids increased; and dihydroferulic acid became doubtful. The 4 indolic acids identified in normal rat urine were unchanged in amount during the first test period with I, but beginning with the 2nd period, indoleacetic (VII) and indolelactic acid (VIII) acids were decreased. III and propionic acid metabolite number 12 disappeared. In the 4th sample 2 new spots appeared, one near VII acid, and the other probably indoleacetylglutamine. In the chromatograms from the 2 patients, there were 24 different urinary phenolic adds identified before I was given, and at first during its administration they all decreased in amount, including Then during the 5th and 6th sampling periods there was a conspicuous increase above the initial normal values for m-hydroxybenzoic, VI, V, and IV acids. II then decreased no further. Of the 13 substances identified on the indole chromatogram, VII and VIII acids decreased in amount, and III increased. The decreased excretions may be explained as due to decarboxylase inhibition by I. The increases in several catabolic products of catechol amine may indicate either an enzyme adaptation or a bypassing of the normal metabolic path. I may effect a secondary

liberation of biogenic amines. 1135-24-6, Cinnamic acid, 4-hydroxy-3-methoxy-IT (in urine, α -methyldopa effect on)

1135-24-6 CAPLUS RN

2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)- (9CI) (CA INDEX NAME) CN

AUTHOR (S):

SOURCE:

1.9 ANSWER 92 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2006:488828 CAPLUS

DOCUMENT NUMBER: 145:431929

TITLE: Combination therapy with tranilast and

angiotensin-converting enzyme inhibition provides additional renoprotection in the remnant kidney model Kelly, D. J.; Zhang, Y.; Cox, A. J.; Gilbert, R. E. Department of Medicine, University of Melbourne, St.

CORPORATE SOURCE: Vincent's Hospital, Victoria, Australia

Kidney International (2006), 69(11), 1954-1960

CODEN: KDYIA5; ISSN: 0085-2538

Nature Publishing Group PUBLISHER:

Journal DOCUMENT TYPE: LANGUAGE: English

Despite current therapy with agents that block the renin-angiotensin system, renal dysfunction continues to progress in a significant proportion of patients with kidney disease. Several pre-clin. studies have reported beneficial effects of tranilast, an inhibitor of transforming growth factor (TGF)- β 's actions in a range of diseases that are characterized by fibrosis. However, whether such therapy provides addnl. benefits in renal disease, when added to angiotensin-converting enzyme (ACE) inhibition, has not been explored. We randomized subtotally (5/6) nephrectomized rats to receive vehicle, the ACE inhibitor, perindopril (6 mg/l), tranilast (400 mg/kg/day), or their combination for 12 wk. When compared with sham-nephrectomized animals, subtotally nephrectomized animals had reduced creatinine clearance, proteinuria, glomerulosclerosis, interstitial fibrosis, tubular atrophy, and evidence of TGF- β activity, as indicated by the abundant nuclear staining of phosphorylated Smad2. These manifestations of injury and $TGF-\beta$ activation were all attenuated by treatment with either tranilast or perindopril, with the latter also attenuating the animals' hypertension. When compared with single-agent treatment, the combination of tranilast and perindopril provided addnl., incremental improvements in creatinine clearance, proteinuria, and glomerulosclerosis, and a reduction in nuclear phsopho-Smad2 beyond single-agent treatment. findings indicate that the combination of tranilast and perindopril was superior to single-agent treatment on kidney structure and function in the remnant kidney model, and suggests the potential for such dual therapy in kidney disease that continues to progress despite blockade of the renin-angiotensin system.

TT 53902-12-8, Tranilast

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

(Biological study); USES (Uses)

(combination of perindopril and tranilast significantly improved creatinine clearance, proteinuria, glomerulosclerosis and reduction in nuclear phsopho-Smad2 compared to perindopril or tranilast alone in subtotally nephrectomized rat)

RN 53902-12-8 CAPLUS

THERE ARE 48 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 48 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

Ь9 ANSWER 93 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

2006:445908 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 144:445358

TITLE: Therapeutic avenanthramide compounds

INVENTOR(S): Meydani, Mohsen

PATENT ASSIGNEE(S): Trustees of Tufts College, USA

SOURCE: U.S. Pat. Appl. Publ., 50 pp., Cont.-in-part of U.S.

> Ser. No. 995,722. CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|-----------|----------|-------------------|------------|
| | | | | |
| US 2006100274 | A1 | 20060511 | US 2005-257918 | 20051025 |
| US 2005239892 | A1 | 20051027 | US 2004-995722 | 20041122 |
| PRIORITY APPLN. INFO.: | | | US 2003-524327P I | 20031121 |
| | | | US 2004-625484P | 20041105 |
| | | | US 2004-995722 A | 2 20041122 |

OTHER SOURCE(S): MARPAT 144:445358

Methods and compns. are disclosed for reducing pro-inflammatory mols., adhesion mols., and vascular smooth muscle cell proliferation, and for increasing NO production The present invention describes the use of phenolic compns., purified from oats or synthetically produced, to decrease the effective amount of pro-inflammatory mols. and/or cell adhesion mols.

Alternatively, an alc. extract or concentrate from oats can be used. The methods

of the present invention can be used as a treatment or prophylaxis of a wide variety of disorders associated with inflammatory states and/or with a lack of or need for nitric oxide (NO), such as inflammatory conditions, pain, free radical associated disorders, cardiovascular diseases, autoimmune disorders, pathol. platelet aggregation, pathol. vasoconstriction, vascular effects of diabetes, stroke, atherosclerosis, hypertension, abnormal vasospasm, and restenosis after angioplasty.

IT 116764-15-9P, Avenanthramide C

RL: NPO (Natural product occurrence); PAC (Pharmacological activity); PNU (Preparation, unclassified); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); USES (Uses)

(therapeutic avenanthramide compds.)

RN116764-15-9 CAPLUS

CNBenzoic acid, 2-[[(2E)-3-(3,4-dihydroxyphenyl)-1-oxo-2-propenyl]amino]-5hydroxy- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

ANSWER 94 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN T.9

2005:1155552 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 143:432646

TITLE: Therapeutic avenanthramide compounds for treatment of

inflammatory-related cardiovascular diseases

PATENT ASSIGNEE(S): Trustees of Tufts College, USA SOURCE:

U.S. Pat. Appl. Publ., 36 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|------------------------|------|----------|-----------------|----|----------|
| | | | | | |
| US 2005239892 | A1 | 20051027 | US 2004-995722 | | 20041122 |
| US 2006100274 | A1 | 20060511 | US 2005-257918 | | 20051025 |
| PRIORITY APPLN. INFO.: | | | US 2003-524327P | P | 20031121 |
| | | | US 2004-625484P | P | 20041105 |
| | | | US 2004-995722 | A2 | 20041122 |

OTHER SOURCE(S): MARPAT 143:432646

GI

AB Methods and compns. are disclosed for reducing pro-inflammatory mols., adhesion mols., and vascular smooth muscle cell proliferation, and for increasing NO production The invention describes the use of phenolic compns. of formula I (R1, R2, or R3 = same or differently H, OH, anhydride, amide,amine, aliphatic, aromatic, acyl, alkoxy, alkylene, alkenylene, alkynylene, hydroxycarbonylalkyl, or heterocyclic group). These compds. which may be purified from oats or synthetically produced decrees the effective amount of pro-inflammatory mols. and/or cell adhesion mols. Alternatively, an alc. extract or concentrate from oats can be used. The methods of the invention

used as a treatment or prophylaxis of a wide variety of disorders associated with inflammatory states and/or with a lack of or need for nitric oxide (NO), such as inflammatory conditions, pain, free radical associated disorders, cardiovascular diseases, autoimmune disorders, pathol. platelet aggregation, pathol. vasoconstriction, vascular effects of diabetes,

stroke, atherosclerosis, hypertension, abnormal vasospasm, and restenosis after angioplasty. Antiproliferative activity of

avenanthramide C (Av-C) in aortic smooth muscle cells was determined as well as Av-C's nitric oxide-inducing effect.

IT 116764-15-9, Avenanthramide C

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

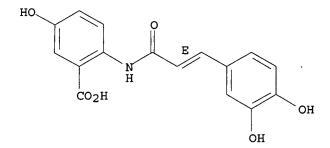
(Biological study); USES (Uses)

(therapeutic avenanthramide compds. for treatment of inflammation-related cardiovascular diseases)

RN116764-15-9 CAPLUS

Benzoic acid, 2-[[(2E)-3-(3,4-dihydroxyphenyl)-1-oxo-2-propenyl]amino]-5-CNhydroxy- (9CI) (CA INDEX NAME)

Double bond geometry as shown.



ANSWER 95 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:700095 CAPLUS

DOCUMENT NUMBER:

143:186499

TITLE:

Tranilast prevents the progression of experimental

diabetic nephropathy through suppression of enhanced

extracellular matrix gene expression

Akahori, Hiroshi; Ota, Tsuguhito; Torita, Muneyoshi; AUTHOR (S):

Ando, Hitoshi; Kaneko, Shuichi; Takamura, Toshinari Department of Diabetes and Digestive Disease, Kanazawa

CORPORATE SOURCE:

University Graduate School of Medical Science,

Ishikawa, Japan

SOURCE: Journal of Pharmacology and Experimental Therapeutics

(2005), 314(2), 514-521

CODEN: JPETAB; ISSN: 0022-3565

PUBLISHER: American Society for Pharmacology and Experimental

Therapeutics

DOCUMENT TYPE:

Journal English

LANGUAGE: The present study was performed to investigate the effects of the antiallergic drug tranilast on the development of diabetic nephropathy in streptozotocin (50 mg/kg)-induced diabetic spontaneously hypertensive rats (SHR). Diabetic SHR were given standard chow or chow containing tranilast at a dose of 1400 mg/kg for 24 wk. The effects of tranilast on urinary albumin excretion, mesangial expansion, expression of transforming growth factor- β (TGF- β) and type I collagen mRNAs, number of anionic sites on the glomerular basement membrane (GBM), and urinary TGF-β and 8-hydroxy-2'-deoxyguanosine (8-OHdG) excretion were assessed. Tranilast did not affect the blood glucose concentration or blood pressure in diabetic SHR. Urinary albumin excretion rate and creatinine clearance were markedly increased in diabetic SHR. Tranilast treatment decreased albuminuria and hyperfiltration. Tranilast inhibited the diabetes-induced expansion of mesangial and tuft areas, as well as the increase in urinary TGF- β and 8-OHdG excretion, loss of anionic sites of GBM, and overexpression of TGF- β as determined immunohistochem. levels of TGF- β and type I collagen mRNA expression were increased in the renal cortex in untreated diabetic SHR at 24 wk, as determined by real-time quant. polymerase chain reaction. Tranilast treatment inhibited the up-regulation of TGF- β and type I collagen mRNA expression by 65 and 36%, resp., in diabetic SHR. In conclusion, tranilast decreased albuminuria by suppressing glomerular hyperfiltration, mesangial expansion, and loss of the charge barrier via regulation of extracellular matrix gene expression and oxidative stress. Tranilast may be clin. useful in the treatment of diabetic nephropathy.

IT 53902-12-8, Tranilast

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(tranilast prevents progression of diabetic nephropathy through suppression of enhanced extracellular matrix gene expression)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

REFERENCE COUNT: 40 THERE ARE 40 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 96 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2003:1009113 CAPLUS

DOCUMENT NUMBER:

141:150692

TITLE:

Intervention with Tranilast Attenuates Renal Pathology

and Albuminuria in Advanced Experimental Diabetic

Nephropathy

AUTHOR(S):

Mifsud, Sally; Kelly, Darren J.; Qi, Weier; Zhang,

Yuan; Pollock, Carol A.; Wilkinson-Berka, Jennifer L.;

Gilbert, Richard E.

CORPORATE SOURCE:

Department of Physiology, University of Melbourne and St. Vincent's Hospital, Melbourne, Vic., Australia

SOURCE:

Nephron (2003), 95(4), p83-p91 CODEN: NPRNAY; ISSN: 0028-2766

PUBLISHER:

S. Karger AG

DOCUMENT TYPE:

Journal

LANGUAGE:

English

Tubulointerstitial pathol. with the accumulation of extracellular matrix are pathol. hallmarks of diabetic nephropathy that are directly related to declining renal function. Tranilast (N-[3,4-dimethoxycinnamoyl]anthranili c acid), an inhibitor of transforming growth factor- β (TGF- β), used to treat hypertrophic scars has recently been shown in pilot studies to exert a beneficial effect in advanced diabetic nephropathy in humans. However, its effects on diabetic renal pathol. are unknown. Studies were conducted using a transgenic model, the diabetic (mRen-2)27 rat, which develops many of the structural and functional characteristics of human diabetic nephropathy when diabetes is induced with streptozotocin (STZ). An exptl. design was chosen to mimic, in part, the clin. context with drug therapy (tranilast 400 mg/kg/day) initiated in established disease (8 wk after STZ) and in the presence of persistent hyperglycemia and hypertension. At 16 wk, diabetes was associated with progressive albuminuria, tubulointerstitial fibrosis and tubular atrophy. Without affecting blood pressure or blood glucose, tranilast treatment was associated with a 83% reduction in tubulointerstitial fibrosis (p < 0.001), a 58% reduction

in tubular atrophy (p < 0.01) and near normalization of albuminuria (p <

0.05) in diabetic Ren-2 rats. In vitro studies in primary cultures of human renal cortical fibroblasts demonstrated a reduction in TGF- β -induced hydroxyproline incorporation and fibronectin synthesis with translast 100 μ M. Translast, when administered during the course of exptl. diabetic nephropathy, attenuates tubulointerstitial pathol. and albuminuria. These findings are consistent with the antagonist effects of translast on TGF- β actions in the diabetic kidney.

IT 53902-12-8, Tranilast

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(tranilast treatment reduced tubulointerstitial fibrosis, tubular atrophy and normalize albuminuria in diabetic rat and also reduced $TGF-\beta$ induced hydroxyproline incorporation and fibronectin synthesis in human renal cortical fibroblasts)

RN 53902-12-8 CAPLUS

CN Benzoic acid, 2-[[3-(3,4-dimethoxyphenyl)-1-oxo-2-propenyl]amino]- (9CI) (CA INDEX NAME)

REFERENCE COUNT: 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 97 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

1997:650742 CAPLUS

DOCUMENT NUMBER:

127:346310

TITLE:

Preparation of tetrahydroquinolinecarboxylic acid

derivatives as intimal thickening inhibitors

INVENTOR(S):

Harada, Hiroshi; Asama, Hiroshi; Nonaka, Yoshiisa;

Kamata, Koji; Hotei, Yukihiko

PATENT ASSIGNEE(S):

Kissei Pharmaceutical Co., Ltd., Japan

SOURCE:

Jpn. Kokai Tokkyo Koho, 6 pp.

CODEN: JKXXAF

DOCUMENT TYPE:

Patent

LANGUAGE:

GI

Japanese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--------|------------|-----------------|----------|
| | | | | |
| JP 09255660 | Α | 19970930 | JP 1996-108763 | 19960325 |
| PRIORITY APPLN. INFO.: | | | JP 1996-108763 | 19960325 |
| OTHER SOURCE(S): | MARPAT | 127:346310 | | |

Ι

$$\begin{array}{c|c}
\text{CO-CH=CH} & & \\
\text{R5} & & & \\
\text{N} & & & \\
\text{CO}_{2}R & & & \\
\text{R3} & & \\
\text{R4} & & & \\
\end{array}$$

AB The title derivs. I [R1 = H, halo, OH, lower alkyl, lower alkoxy, cycloalkylalkoxy, aralkyloxy, lower acyl, mono- or di(lower alkyl)amino, lower alkoxycarbonyl; R2-3 = H, halo, lower alkyl, lower alkoxy, cycloalkylalkoxy, aralkyloxy; R4-6 = H, OH, lower alkyl, lower alkoxy, CO2H, lower alkoxycarbonyl; R = H, lower alkyl] and their pharmacol. acceptable salts are claimed. I inhibit hyperproliferation of intimal cells to prevent atherosclerosis and restenosis after PTCA and DCA (directional coronary atherectomy). 1-(3,4,5-Trimethoxycinnamoyl)-1,2,3,4-tetrahydroquinoline-2-carboxylic acid (preparation given) inhibited proliferation of smooth muscle cells of thoracic aorta isolated from a spontaneously hypertensive rat at IC50 32 μM.

IT 90-50-6, 3,4,5-Trimethoxycinnamic acid

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of N-cinnamoyltetrahydroquinolinecarboxylic acid derivs. as intimal thickening inhibitors)

RN 90-50-6 CAPLUS

CN 2-Propenoic acid, 3-(3,4,5-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

L9 ANSWER 98 OF 98 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1997:648534 CAPLUS

DOCUMENT NUMBER: 127:346309

TITLE: Preparation of decahydroquinolinecarboxylic acid

derivatives as intimal thickening inhibitors

INVENTOR(S): Harada, Hiroshi; Kusama, Hiroshi; Nonaka, Yoshiisa;

Kamata, Koji; Hotei, Yukihiko

PATENT ASSIGNEE(S): Kissei Pharmaceutical Co., Ltd., Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 5 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--------|------------|-----------------|----------|
| | | | | |
| JP 09255661 | A | 19970930 | JP.1996-108764 | 19960325 |
| PRIORITY APPLN. INFO.: | | | JP 1996-108764 | 19960325 |
| OTHER SOURCE(S): | MARPAT | 127:346309 | | |
| GI | | | | |

$$\begin{array}{c|c}
\text{CO-CH=CH} & & \\
\text{R5} & & \\
\text{R6} & & \\
\text{R4} & & \\
\end{array}$$

AB The title derivs. I [R1 = H, halo, OH, lower alkyl, lower alkoxy,

Ι

cycloalkylalkoxy, aralkyloxy, lower acyl, mono- or di(lower alkyl)amino, lower alkoxycarbonyl; R2-3 = H, halo, lower alkyl, lower alkoxy, cycloalkylalkoxy, aralkyloxy; R4-6 = H, OH, lower alkyl, lower alkoxy, CO2H, lower alkoxycarbonyl; R = H, lower alkyl] and their pharmacol. acceptable salts are claimed. I inhibit hyperproliferation of intimal cells to prevent atherosclerosis and restenosis after PTCA and DCA (directional coronary atherectomy). 1-(3,4,5-Trimethoxycinnamoyl)decahydroquinoline-2-carboxylic acid (preparation given) inhibited proliferation of smooth muscle cells of thoracic aorta isolated from a spontaneously hypertensive rat at IC50 104 μ M. 90-50-6, 3,4,5-Trimethoxycinnamic acid RL: RCT (Reactant); RACT (Reactant or reagent)

L: RCT (Reactant); RACT (Reactant or reagent)
(preparation of N-cinnamoyldecahydroquinolinecarboxylic acid derivs. as
intimal thickening inhibitors)

RN 90-50-6 CAPLUS

IT

CN 2-Propenoic acid, 3-(3,4,5-trimethoxyphenyl)- (9CI) (CA INDEX NAME)

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L2
     ANSWER 102 OF 102 REGISTRY COPYRIGHT 2007 ACS on STN
RN
     77-95-2 REGISTRY
ED
     Entered STN: 16 Nov 1984
     Cyclohexanecarboxylic acid, 1,3,4,5-tetrahydroxy-,
CN
     (1\alpha, 3R, 4\alpha, 5R) - (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Cyclohexanecarboxylic acid, 1,3,4,5-tetrahydroxy-, (-)- (8CI)
CN
     Cyclohexanecarboxylic acid, 1,3,4,5-tetrahydroxy-, [1R-
CN
     (1\alpha, 3\alpha, 4\alpha, 5\beta)] -
OTHER NAMES:
CN
     (-)-Quinic acid
     D-(-)-Quinic acid
CN
CN
     D-Quinic acid
CN
     Quinic acid
     Quinic acid, (-)-
CN
     STEREOSEARCH
FS
DR
     35949-55-4
MF
     C7 H12 O6
     COM
CI
                  AGRICOLA, ANABSTR, BEILSTEIN*, BIOSIS, BIOTECHNO, CA, CABA,
LC
     STN Files:
       CAOLD, CAPLUS, CASREACT, CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN, CSCHEM,
       EMBASE, GMELIN*, IFICDB, IFIUDB, IPA, MEDLINE, MRCK*, NAPRALERT, PROMT,
       PS, RTECS*, SYNTHLINE, TOXCENTER, USPAT2, USPATFULL
         (*File contains numerically searchable property data)
     Other Sources: DSL**, EINECS**
         (**Enter CHEMLIST File for up-to-date regulatory information)
```

Absolute stereochemistry. Rotation (-).

=>

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

771 REFERENCES IN FILE CA (1907 TO DATE)

46 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

774 REFERENCES IN FILE CAPLUS (1907 TO DATE)

17 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

```
L3
     ANSWER 120 OF 127 REGISTRY COPYRIGHT 2007 ACS on STN
     1135-24-6 REGISTRY
RN
ED
     Entered STN: 16 Nov 1984
     2-Propenoic acid, 3-(4-hydroxy-3-methoxyphenyl)- (9CI)
                                                              (CA INDEX NAME)
CN
OTHER CA INDEX NAMES:
     Cinnamic acid, 4-hydroxy-3-methoxy- (7CI, 8CI)
OTHER NAMES:
     3-(4-Hydroxy-3-methoxyphenyl)-2-propenoic acid
CN
     3-(4-Hydroxy-3-methoxyphenyl)acrylic acid
CN
     3-Methoxy-4-hydroxycinnamic acid
CN
     4'-Hydroxy-3'-methoxycinnamic acid
CN
     4-Hydroxy-3-methoxycinnamic acid
CN
     Coniferic acid
CN
     Ferulaic acid
CN
     Ferulic acid
CN
     NSC 2821
CN
     NSC 51986
CN
CN
     NSC 674320
MF
     C10 H10 O4
CI
     COM
LC
                  AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN*, BIOSIS, BIOTECHNO, CA,
     STN Files:
       CABA, CAOLD, CAPLUS, CASREACT, CBNB, CHEMCATS, CHEMINFORMRX, CHEMLIST,
       CIN, CSCHEM, DDFU, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE,
       MRCK*, NAPRALERT, PIRA, PROMT, RTECS*, SPECINFO, SYNTHLINE, TOXCENTER,
       ULIDAT, USPAT2, USPATFULL, VETU
         (*File contains numerically searchable property data)
     Other Sources:
                      EINECS**, NDSL**, TSCA**
         (**Enter CHEMLIST File for up-to-date regulatory information)
```

$$\begin{array}{c} \text{CH-CO}_2\text{H} \\ \\ \text{OMe} \end{array}$$

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

7378 REFERENCES IN FILE CA (1907 TO DATE)
444 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
7443 REFERENCES IN FILE CAPLUS (1907 TO DATE)
2 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

```
ANSWER 45 OF 45 REGISTRY COPYRIGHT 2007 ACS on STN
L4
RN
     327-97-9 REGISTRY
     Entered STN: 16 Nov 1984
ED
     Cyclohexanecarboxylic acid, 3-[[3-(3,4-dihydroxyphenyl)-1-oxo-2-
CN
     propenyl]oxy]-1,4,5-trihydroxy-, (1S,3R,4R,5R)- (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Chlorogenic acid (8CI)
CN
     Cyclohexanecarboxylic acid, 3-[[3-(3,4-dihydroxyphenyl)-1-oxo-2-
CN
     propenyl] \alpha oxy] -1,4,5-trihydroxy-, [1S-(1\alpha,3\beta,4\alpha,5\alpha)] -
OTHER NAMES:
     3-(3,4-Dihydroxycinnamoyl)quinic acid
CN
     3-Caffeoylquinic acid
CN
CN
     3-O-(3,4-Dihydroxycinnamoyl)-D-quinic acid
CN
     3-O-Caffeoylquinic acid
CN
     Heriguard
CN
     NSC 407296
     NSC 70861
CN
FS
     STEREOSEARCH
     12626-41-4, 15076-00-3, 16310-14-8, 16431-25-7, 16431-26-8, 108657-60-9
DR
MF
     C16 H18 O9
CI
     COM
LC
     STN Files:
                  AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN*, BIOSIS, BIOTECHNO, CA,
       CABA, CAOLD, CAPLUS, CASREACT, CBNB, CHEMCATS, CHEMLIST, CIN, CSCHEM,
       DDFU, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*,
       MSDS-OHS, NAPRALERT, PIRA, PROMT, RTECS*, SPECINFO, TOXCENTER, USPAT2,
       USPATFULL, VETU
         (*File contains numerically searchable property data)
     Other Sources:
                      EINECS**
         (**Enter CHEMLIST File for up-to-date regulatory information)
```

Absolute stereochemistry.

Double bond geometry unknown.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

6187 REFERENCES IN FILE CA (1907 TO DATE)
247 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
6236 REFERENCES IN FILE CAPLUS (1907 TO DATE)
32 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

```
ANSWER 1 OF 1 REGISTRY COPYRIGHT 2007 ACS on STN
L5
     104594-70-9 REGISTRY
RN
     Entered STN: 11 Oct 1986
ED
     2-Propenoic acid, 3-(3,4-dihydroxyphenyl)-, 2-phenylethyl ester (9CI)
CN
     INDEX NAME)
OTHER NAMES:
     \beta-Phenylethyl caffeate
CN
     2-Phenylethyl caffeate
CN
     Caffeic acid phenethyl ester
CN
     132031-37-9
DR
     C17 H16 O4
MF
SR
     CA
                  ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOSIS, BIOTECHNO,
LC
     STN Files:
       CA, CAPLUS, CASREACT, CHEMCATS, CSCHEM, EMBASE, IPA, MEDLINE, RTECS*,
       TOXCENTER, USPAT7, USPATFULL
         (*File contains numerically searchable property data)
```

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

313 REFERENCES IN FILE CA (1907 TO DATE)

4 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

317 REFERENCES IN FILE CAPLUS (1907 TO DATE)

```
L6 ANSWER 8 OF 8 REGISTRY COPYRIGHT 2007 ACS on STN
```

RN 20283-92-5 REGISTRY

ED Entered STN: 16 Nov 1984

CN Benzenepropanoic acid, α -[[(2E)-3-(3,4-dihydroxyphenyl)-1-oxo-2-propenyl]oxy]-3,4-dihydroxy-, (α R)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Benzenepropanoic acid, α -[[3-(3,4-dihydroxyphenyl)-1-oxo-2-propenyl]oxy]-3,4-dihydroxy-, [R-(E)]-

CN Cinnamic acid, 3,4-dihydroxy-, 2-ester with 3-(3,4-dihydroxyphenyl)lactic acid (8CI)

CN Rosmarinic acid (6CI, 7CI)

OTHER NAMES:

CN Mamorekku RUH 2

CN RM 21A

CN Rosemaric acid

CN Rosemary acid

FS STEREOSEARCH

MF C18 H16 O8

CI COM

LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN*, BIOSIS, BIOTECHNO, CA, CAOLD, CAPLUS, CASREACT, CBNB, CHEMCATS, CHEMLIST, CIN, CSCHEM, DDFU, DRUGU, EMBASE, IPA, NAPRALERT, PHAR, PROMT, PROUSDDR, RTECS*, SPECINFO, TOXCENTER, USPAT2, USPATFULL (*File contains numerically searchable property data)

Absolute stereochemistry. Rotation (+).

Double bond geometry as shown.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

928 REFERENCES IN FILE CA (1907 TO DATE)

18 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

939 REFERENCES IN FILE CAPLUS (1907 TO DATE)

5 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

```
L7
     ANSWER 17 OF 17 REGISTRY COPYRIGHT 2007 ACS on STN
     469-36-3 REGISTRY
RN
     Entered STN: 16 Nov 1984
ED
     9,19-Cyclolanostan-3-ol, 24-methylene-, 3-(4-hydroxy-3-methoxyphenyl)-2-
CN
     propenoate, (3β) - (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     1H,19H-Cyclopropa[9,10]cyclopenta[a]phenanthrene, 9,19-cyclolanostan-3-ol
CN
     deriv.
     9,19-Cyclo-9\beta-lanostan-3\beta-ol, 24-methylene-,
CN
     4-hydroxy-3-methoxycinnamate (8CI)
     Cinnamic acid, 4-hydroxy-3-methoxy-, 24-methylene-9,19-cyclo-9β-
CN
     lanostan-3\beta-yl ester (8CI)
     Oryzanol C (6CI)
CN
OTHER NAMES:
     24-Methylenecycloartanol ferulate
CN
     24-Methylenecycloartanol ferulic acid ester
CN
     24-Methylenecycloartanyl ferulate
CN
     STEREOSEARCH
FS
     97818-51-4, 119104-04-0, 100109-90-8, 116846-86-7
DR
MF
     C41 H60 O4
CI
     COM
                  AGRICOLA, BEILSTEIN*, BIOSIS, CA, CAOLD, CAPLUS, CHEMCATS,
LC
     STN Files:
       MRCK*, TOXCENTER, USPATFULL
```

(*File contains numerically searchable property data)

Absolute stereochemistry.

Double bond geometry unknown.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

55 REFERENCES IN FILE CA (1907 TO DATE)
56 REFERENCES IN FILE CAPLUS (1907 TO DATE)
3 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

=>

```
L8
     ANSWER 66 OF 66 REGISTRY COPYRIGHT 2007 ACS on STN
     458-37-7 REGISTRY
RN
     Entered STN: 16 Nov 1984
ED
     1,6-Heptadiene-3,5-dione, 1,7-bis(4-hydroxy-3-methoxyphenyl)-, (1E,6E)-
CN
           (CA INDEX NAME)
     (9CI)
OTHER CA INDEX NAMES:
     1,6-Heptadiene-3,5-dione, 1,7-bis(4-hydroxy-3-methoxyphenyl)-, (E,E)-
     (8CI)
     Curcumin (6CI)
CN
OTHER NAMES:
     (E,E)-1,7-Bis(4-hydroxy-3-methoxyphenyl)-1,6-heptadiene-3,5-dione
CN
     C Yellow 15
     C.I. 75300
CN
     C.I. Natural Yellow 3
CN
     Curcuma
CN
     Curcumin I
CN
     Curcumine
CN
     Diferuloylmethane
CN
     E 100
CN
     E 100 (dye)
CN
CN
     Haidr
CN
     Halad
     Haldar
CN
CN
     Halud
     Indian Saffron
CN
     Kacha Haldi
CN
CN
     Merita Earth
     Natural Yellow 3
CN
     NSC 32982
CN
     San-Ei Curcumine AL
CN
     San-Ei Gen Curcumine AL
CN
CN
     Souchet
     Terra Merita
CN
     trans, trans-Curcumin
CN
     Turmeric
CN
CN
     Turmeric (dye)
CN
     Turmeric yellow
CN
     Ukon
CN
     Ukon (dye)
CN
     Yellow Ginger
CN
     Yellow Root
CN
     Yo-Kin
FS
     STEREOSEARCH
DR
     15845-47-3, 73729-23-4, 79257-48-0, 91884-86-5, 33171-04-9
MF
     C21 H20 O6
CI
LC
     STN Files:
                  ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN*,
       BIOSIS, BIOTECHNO, CA, CABA, CAOLD, CAPLUS, CASREACT, CBNB, CHEMCATS,
       CHEMLIST, CIN, CSCHEM, DDFU, DRUGU, EMBASE, HSDB*, IFICDB, IFIPAT,
       IFIUDB, IMSRESEARCH, IPA, MEDLINE, MRCK*, MSDS-OHS, NAPRALERT, PHAR,
       PIRA, PROMT, PROUSDDR, RTECS*, SPECINFO, SYNTHLINE, TOXCENTER, ULIDAT,
       USPAT2, USPATFULL
         (*File contains numerically searchable property data)
                      DSL**, EINECS**, TSCA**
     Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
```

Double bond geometry as shown.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

2843 REFERENCES IN FILE CA (1907 TO DATE)

132 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

2868 REFERENCES IN FILE CAPLUS (1907 TO DATE)

21 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

=>

```
ANSWER 116 OF 118 REGISTRY COPYRIGHT 2007 ACS on STN
L9
     331-39-5 REGISTRY
RN
     Entered STN: 16 Nov 1984
ED
     2-Propenoic acid, 3-(3,4-dihydroxyphenyl)- (9CI)
                                                        (CA INDEX NAME)
CN
OTHER CA INDEX NAMES:
     Cinnamic acid, 3,4-dihydroxy- (8CI)
OTHER NAMES:
     3,4-Dihydroxybenzeneacrylic acid
CN
     3,4-Dihydroxycinnamic acid
CN
     3-(3,4-Dihydroxyphenyl)-2-propenoic acid
CN
     3-(3,4-Dihydroxyphenyl)propenoic acid
CN
     4-(2'-Carboxyvinyl)-1,2-dihydroxybenzene
CN
     4-(2-Carboxyethenyl)-1,2-dihydroxybenzene
CN
CN
     Caffeic acid
     NSC 57197
CN
CN
     NSC 623438 .
     C9 H8 O4
MF
CI
     COM
LC
                  ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN*, BIOSIS,
     STN Files:
       BIOTECHNO, CA, CABA, CAOLD, CAPLUS, CASREACT, CHEMCATS, CHEMLIST, CIN,
       CSCHEM, DDFU, DRUGU, EMBASE, HSDB*, IFICDB, IFIPAT, IFIUDB, IPA,
       MEDLINE, MRCK*, MSDS-OHS, NAPRALERT, PIRA, PROMT, PS, RTECS*, SPECINFO,
       SYNTHLINE, TOXCENTER, ULIDAT, USPATZ, USPATFULL, VETU
         (*File contains numerically searchable property data)
                     DSL**, EINECS**
     Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
```

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

7275 REFERENCES IN FILE CA (1907 TO DATE)

423 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

7329 REFERENCES IN FILE CAPLUS (1907 TO DATE)

12 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

```
ANSWER 66 OF 66 REGISTRY COPYRIGHT 2007 ACS on STN
L19
     458-37-7 REGISTRY
RN
     Entered STN: 16 Nov 1984
ED
     1,6-Heptadiene-3,5-dione, 1,7-bis(4-hydroxy-3-methoxyphenyl)-, (1E,6E)-
CN
            (CA INDEX NAME)
     (9CI)
OTHER CA INDEX NAMES:
     1,6-Heptadiene-3,5-dione, 1,7-bis(4-hydroxy-3-methoxyphenyl)-, (E,E)-
     (8CI)
     Curcumin (6CI)
CN
OTHER NAMES:
     (E,E)-1,7-Bis(4-hydroxy-3-methoxyphenyl)-1,6-heptadiene-3,5-dione
CN
     C Yellow 15
CN
     C.I. 75300
CN
     C.I. Natural Yellow 3
CN
     Curcuma
CN
     Curcumin I
CN
     Curcumine
CN
     Diferuloylmethane
CN
CN
     E 100
     E 100 (dye)
CN
     Haidr
CN
CN
     Halad
     Haldar
CN
     Halud
CN
CN
     Indian Saffron
     Kacha Haldi
CN
CN
     Merita Earth
     Natural Yellow 3
CN
     NSC 32982
CN
     San-Ei Curcumine AL
CN
     San-Ei Gen Curcumine AL
CN
CN
     Souchet
CN
     Terra Merita
CN
     trans, trans-Curcumin
CN
     Turmeric
CN
     Turmeric (dye)
CN
     Turmeric yellow
CN
     Ukon
CN
     Ukon (dye)
CN
     Yellow Ginger
CN
     Yellow Root
CN
     Yo-Kin
FS
     STEREOSEARCH
     15845-47-3, 73729-23-4, 79257-48-0, 91884-86-5, 33171-04-9
DR
MF
     C21 H20 O6
CI
LC
     STN Files:
                  ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN*,
       BIOSIS, BIOTECHNO, CA, CABA, CAOLD, CAPLUS, CASREACT, CBNB, CHEMCATS,
       CHEMLIST, CIN, CSCHEM, DDFU, DRUGU, EMBASE, HSDB*, IFICDB, IFIPAT,
       IFIUDB, IMSRESEARCH, IPA, MEDLINE, MRCK*, MSDS-OHS, NAPRALERT, PHAR,
       PIRA, PROMT, PROUSDDR, RTECS*, SPECINFO, SYNTHLINE, TOXCENTER, ULIDAT,
       USPAT2, USPATFULL
         (*File contains numerically searchable property data)
                      DSL**, EINECS**, TSCA**
     Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
```

Double bond geometry as shown.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

=>

2843 REFERENCES IN FILE CA (1907 TO DATE)

132 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

2868 REFERENCES IN FILE CAPLUS (1907 TO DATE)

21 REFERENCES IN FILE CAOLD (PRIOR TO 1967)